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(54) Title: HUMAN PROTEINS HAVING HYDROPHOBIC DOMAINS AND DNAS ENCODING THESE PROTEINS

(57) Abstract: The present invention provides human proteins having hydrophobic domains, DNAs encoding these proteins, expression vectors for these DNAs, transformed eukaryotic cells expressing these DNAs and antibodies directed to these proteins.





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DESCRIPTION

Human Proteins Having Hydrophobic

Domains and DNAs Encoding These Proteins

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TECHNICAL FIELD

The present invention relates to human proteins having hydrophobic domains, DNAs encoding these proteins, expression vectors for these DNAs, eukaryotic cells expressing these DNAs and antibodies directed to these proteins. The proteins of the present invention can be employed as pharmaceuticals or as antigens for preparing antibodies directed to these proteins. The human cDNAs of the present invention can be utilized as probes for genetic diagnosis and gene sources for gene therapy. Furthermore, the cDNAs can be utilized as gene sources for producing the proteins encoded by these cDNAs in large quantities. Cells into which these genes are introduced to express secretory proteins or membrane proteins in large quantities can be utilized for detection of the corresponding receptors or ligands, screening of novel small molecule pharmaceuticals and the like. The antibodies of the present invention can be utilized for the detection, quantification, purification and the like of the proteins of the present invention.

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BACKGROUND ART

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Cells secrete many proteins extracellularly. These secretory proteins play important roles in the proliferation control, the differentiation induction, the material transport, the biophylaxis, and the like of the cells. Unlike intracellular proteins, the secretory proteins exert their actions outside the cells. Therefore, they can be administered in the intracorporeal manner such as the injection or the drip, so that they possess hidden potentialities as pharmaceuticals. In fact, a number of human secretory proteins such as interferons, interleukins, erythropoietin, thrombolytic agents and the like are pharmaceuticals. currently employed as In addition, secretory proteins other than those described above are undergoing clinical trials for developing their use as pharmaceuticals. It is believed that the human cells produce many unknown secretory proteins. Availability of these secretory proteins as well as genes encoding them is expected to lead to development of novel pharmaceuticals utilizing them.

On the other hand, membrane proteins play important roles, as signal receptors, ion channels, transporters and the like in the material transport and the signal transduction through the cell membrane. Examples thereof include receptors for various cytokines, ion

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channels for the sodium ion, the potassium ion, the chloride ion and the like, transporters for saccharides and amino acids and the like. The genes for many of them have already been cloned. It has been clarified that abnormalities in these membrane proteins are involved in a number of previously cryptogenic diseases. Therefore, discovery of a new membrane protein is expected to lead to elucidation of the causes of many diseases, so that isolation of new genes encoding the membrane proteins has been desired.

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Heretofore, due to difficulty in the purification from human cells, many of these secretory proteins and membrane proteins have been isolated by genetic approaches. A general method is the so-called expression cloning method, in which a cDNA library is introduced into eukaryotic cells to express cDNAs, and the cells secreting, or expressing on the surface of membrane, the protein having the activity of interest are then screened. However, only genes for proteins with known functions can be cloned by using this method.

In general, a secretory protein or a membrane protein possesses at least one hydrophobic domain within the protein. After synthesis on ribosomes, such domain works as a secretory signal or remains in the phospholipid membrane to be entrapped in the membrane. Accordingly, if the existence of a highly hydrophobic domain is observed in the amino acid sequence of a protein encoded by a cDNA when the

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whole base sequence of the full-length cDNA is determined, it is considered that the cDNA encodes a secretory protein or a membrane protein.

5 OBJECTS OF INVENTION

The main object of the present invention is to provide novel human proteins having hydrophobic domains, DNAs encoding these proteins, expression vectors for these DNAs, transformed eukaryotic cells that are capable of expressing these DNAs and antibodies directed to these proteins. This object as well as other objects and advantages of the present invention will become apparent to those skilled in the art from the following description with reference to the accompanying drawings.

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SUMMARY OF INVENTION

As the result of intensive studies, the present inventors have successfully cloned cDNAs encoding proteins having hydrophobic domains from the human full-length cDNA bank, thereby completing the present invention. Thus, the present invention provides a human protein having hydrophobic domain(s), namely a protein comprising any one of an amino acid sequence selected from the group consisting of SEQ ID NOS: 1 to 10, 31 to 40, 61 to 70, 91 to 100 and 121 to 130. Moreover, the present invention provides a DNA

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encoding said protein, exemplified by a cDNA comprising any one of a base sequence selected from the group consisting of SEQ ID NOS: 11 to 30, 41 to 60, 71 to 90, 101 to 120 and 131 to 150, an expression vector that is capable of expressing said DNA by in vitro translation or in eukaryotic cells, a transformed eukaryotic cell that is capable of expressing said DNA and of producing said protein and an antibody directed to said protein.

10 BRIEF DESCRIPTION OF DRAWINGS

Fig. 1 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03171.

Fig. 2 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03424.

Fig. 3 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03444.

20 Fig. 4 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03478.

Fig. 5 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03499.

		Fig.	6	il	llus	trat	es	the
	hydrophob	icity/hydr	ophilicity	profile	of	the	protein	encoded
	by clone	нр03500.						
		Fig.	7	i	llus	strat	es	the
5	hydrophob	icity/hydr	ophilicity	profile	of	the	protein	encoded
	by clone	нр10691.						
		Fig.	8	i	llus	strat	es	the
	hydrophob	oicity/hydr	ophilicity	profile	of	the	protein	encoded
	by clone	HP10703.						
10		Fig.	9	i	llus	strat	tes	the
	hydrophob	oicity/hydr	ophilicity	profile	of	the	protein	encoded
	by clone	HP10711.						
		Fig.	. 10	i	llu	stra	tes	the
	hydrophob	oicity/hydr	ophilicity	profile	of	the	protein	encoded
15	by clone	HP10712.						
		Fig.	11	i	llu	stra	tes	the
	hydrophol	oicity/hydr	ophilicity	profile	of	the	protein	encoded
	by clone	нр03010.						
		Fig.	. 12	i	illu	stra	tes	the
20	hydrophol	oicity/hydr	cophilicity	profile	of	the	protein	encoded
	by clone	нр03576.	-					
		Fig.	13	:	illı	ıstra	ites	the
	hydrophol	bicity/hyd:	rophilicity	profile	of	the	protein	encoded
	by clone	нр03611.						
25		Fig.	14	•	illı	ıstra	ates	the

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hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03612.

Fig. 15 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10407.

Fig. 16 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10713.

Fig. 17 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10714.

Fig. 18 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10716.

15 Fig. 19 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10717.

Fig. 20 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10718.

Fig. 21 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP03745.

Fig. 22 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded

by clone HP03747.

Fig. 23 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded

by clone HP10719.

by clone HP10/19.

5 Fig. 24 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10720.

Fig. 25 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded

10 by clone HP10721.

Fig. 26 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10725.

Fig. 27 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10727.

Fig. 28 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10728.

20 Fig. 29 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10730.

Fig. 30 illustrates the

hydrophobicity/hydrophilicity profile of the protein encoded

25 by clone HP10742.

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		Fig.	31	i	llu	strat	tes .	the
•	hydrophol	bicity/hyd	rophilicity	profile	of	the	protein	encoded
	by clone	HP03800.				•	•	
		Fig.	32	i	llus	strat	tes	the
5	hydrophol	bicity/hyd	rophilicity	profile	of	the	protein	encoded
	by clone	HP03831.						
		Fig.	33	i	llu	strat	tes	the
	hydrophol	bicity/hyd	rophilicity	profile	of	the	protein	encoded
•	by clone	нр03879.						
10		Fig.	34	i	llu	strat	tes	the
	hydropho	bicity/hyd	rophilicity	profile	of	the	protein	encoded
	by clone	нр03880.						•
		Fig.	35	i	llu	stra	tes	the
	hydropho	bicity/hyd	rophilicity	profile	of	the	protein	encoded
15	by clone	HP10704.						
		Fig.	36	i	llu	stra	tes	the
	hydropho	bicity/hyd	rophilicity	profile	of	the	protein	encoded
	by clone	нр10715.						
•	•	Fig.	37	·	llu	stra [.]	tes	the
20	hydropho	bicity/hyd	lrophilicity	profile	of	the	protein	encoded
	by clone	HP10724.						
		Fig.	38	i	llu	stra	tes	the
	hydropho	bicity/hyd	lrophilicity	profile	of	the	protein	encoded
	by clone	HP10733.						
25		Fig.	39	i	llu	stra	tes	the

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Fig.

hydrophol	oicity/hydro	philicity	profile	of	the	protein	encoded
by clone	HP10734.						
	Fig.	40	i	llu	stra	tes	the
hydrophol	oicity/hydro	philicity	profile	of	the	protein	encoded
by clone	HP10756.						
	Fig.	41	i	llu	stra	tes	the
hydrophol	oicity/hydro	philicity	profile	of	the	protein	encoded
by clone	нр03670.						
	Fig.	42	i	llu	stra	tes	the
hydrophol	bicity/hydro	philicity	profile	of	the	protein	encoded
by clone	HP03688.						
	Fig.	43	i	llu	stra [.]	tes	the
hydrophol	bicity/hydro	philicity	profile	of	the	protein	encoded
by clone	HP03825						
	Fig.	44	i	llu	stra	tes	the
hydrophol	bicity/hydro	philicity	profile	of	the	protein	encoded
by clone	нр03877.						
	Fig.	45	i	llu	stra	tes	the
hydrophol	bicity/hydro	philicity	profile	of	the	protein	encoded
by clone	HP10765.						
	Fig.	. 46	i	llu	stra	tes	the
hydropho	bicity/hydro	philicity	profile	of	the	protein	encoded
by clone	HP10766.						

47 illustrates

hydrophobicity/hydrophilicity profile of the protein encoded

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by clone HP10770.

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Fig. 48 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10772.

Fig. 49 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10773.

Fig. 50 illustrates the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10776.

DETAILED DESCRIPTION OF THE INVENTION

The proteins of the present invention can be obtained, for example, by a method for isolating proteins from human organs, cell lines or the like, a method for 15 preparing peptides by the chemical synthesis based on the amino acid sequences of the present invention, or a method for producing proteins by the recombinant DNA technology using the DNAs encoding the hydrophobic domains of the present invention. Among these, the method for producing 20 proteins by the recombinant DNA technology is preferably employed. For example, the proteins can be expressed in vitro by preparing an RNA by in vitro transcription from a vector having the cDNA of the present invention, and then carrying out in vitro translation using this RNA as a 25

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template. Alternatively, incorporation of the translated region into a suitable expression vector by the method known in the art may lead to expression of a large amount of the encoded protein in prokaryotic cells such as *Escherichia coli*, *Bacillus subtilis*, etc., and eukaryotic cells such as veasts, insect cells, mammalian cells, etc.

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In the case where the protein of the present invention is produced by expressing the DNA by in vitro translation, the protein of the present invention can be produced in vitro by incorporating the translated region of this cDNA into a vector having an RNA polymerase promoter, and then adding the vector to an in vitro translation system such as a rabbit reticulocyte lysate or a wheat germ extract, which contains an RNA polymerase corresponding to the promoter. The RNA polymerase promoters are exemplified by T7, T3, SP6 and the like. The vectors containing promoters for these RNA polymerases are exemplified by pKA1, pCDM8, pT3/T7 18, pT7/3 19, pBluescript II and the like. Furthermore, the protein of the present invention can be expressed in the secreted form or the form incorporated in the microsome membrane when a canine pancreas microsome or the like is added to the reaction system.

In the case where the protein of the present invention is produced by expressing the DNA in a microorganism such as Escherichia coli etc., a recombinant

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expression vector in which the translated region of the cDNA of the present invention is incorporated into an expression vector having an origin which is capable of replicating in the microorganism, a promoter, a ribosome-binding site, a cDNA-cloning site, a terminator and the like is constructed. After transformation of the host cells with this expression vector, the resulting transformant is cultivated, whereby the protein encoded by the cDNA can be produced in large quantities in the microorganism. In this case, a protein fragment containing any translated region can be obtained by adding an initiation codon and a termination codon in front of and behind the selected translated region to express the protein. Alternatively, the protein can be expressed as a fusion protein with another protein. Only the portion of the protein encoded by the cDNA can be obtained by cleaving this fusion protein with a suitable protease. The expression vectors for Escherichia coli are exemplified by the pUC series, pBluescript II, the pET expression system, the pGEX expression system and the like.

In the case where the protein of the present invention is produced by expressing the DNA in eukaryotic cells, the protein of the present invention can be produced as a secretory protein, or as a membrane protein on the surface of cell membrane, by incorporating the translated region of the cDNA into an expression vector for eukaryotic 25

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cells that has a promoter, a splicing region, a poly(A) addition site and the like, and then introducing the vector into the eukaryotic cells. The expression vectors are exemplified by pKA1, pED6dpc2, pCDM8, pSVK3, pMSG, pSVL, pBK-CMV, pBK-RSV, EBV vectors, pRS, pYES2 and the like. Examples of eukaryotic cells to be used in general include mammalian cultured cells such as monkey kidney COS7 cells, Chinese hamster ovary CHO cells and the like, budding yeasts, fission yeasts, silkworm cells, Xenopus oocytes and the like. Any eukaryotic cells may be used as long as they are capable of expressing the proteins of the present invention. The expression vector can be introduced into the eukaryotic cells by using a method known in the art such as the electroporation method, the calcium phosphate method, the liposome method, the DEAE-dextran method and the like.

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After the protein of the present invention is expressed in prokaryotic cells or eukaryotic cells, the protein of interest can be isolated and purified from the culture by a combination of separation procedures known in the art. Examples of the separation procedures include treatment with a denaturing agent such as urea or a detergent, sonication, enzymatic digestion, salting-out or precipitation, solvent dialysis, centrifugation, ultrafiltration, gel filtration, SDS-PAGE, isoelectric focusing, ion-exchange chromatography, hydrophobic

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chromatography, affinity chromatography, reverse phase chromatography and the like.

The proteins of the present invention also include peptide fragments (of 5 amino acid residues or more) containing any partial amino acid sequences in the amino acid sequences represented by SEQ ID NOS: 1 to 10, 31 to 40, 61 to 70, 91 to 100 and 121 to 130. These peptide fragments can be utilized as antigens for preparation of antibodies. Among the proteins of the present invention, those having the signal sequences are secreted in the form of mature proteins after the signal sequences are removed. Therefore, these mature proteins shall come within the scope of the protein of the present invention. The N-terminal amino acid sequences of the mature proteins can be easily determined by using the method for the determination of cleavage site of a signal sequence [JP-A 8-187100]. Furthermore, some membrane proteins undergo the processing on the cell surface to be converted to the secreted forms. Such proteins or peptides in the secreted forms shall also come within the scope of the protein of the present invention. In the case where sugar chain-binding sites are present in the amino acid sequences of the proteins, expression of the proteins in appropriate eukaryotic cells affords the proteins to which sugar chains are added. Accordingly, such proteins or peptides to which sugar chains are added shall also come

within the scope of the protein of the present invention.

The DNAs of the present invention include all the DNAs encoding the above-mentioned proteins. These DNAs can be obtained by using a method for chemical synthesis, a method for cDNA cloning and the like.

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The cDNAs of the present invention can be cloned, for example, from cDNA libraries derived from the human cells. The cDNAs are synthesized by using poly(A)* RNAs extracted from human cells as templates. The human cells may be cells delivered from the human body, for example, by the operation or may be the cultured cells. The cDNAs can be synthesized by using any method such as the Okayama-Berg method [Okayama, H. and Berg, P., Mol. Cell. Biol. 2: 161-170 (1982)], the Gubler-Hoffman method [Gubler, U. and Hoffman, J., Gene 25: 263-269 (1983)] and the like. However, it is desirable to use the capping method [Kato, S. et al., Gene 150: 243-250 (1994)], as exemplified in Examples, in order to obtain a full-length clone in an effective manner. In addition, commercially available human cDNA libraries can be utilized. The cDNAs of the present invention can be cloned libraries from the CDNA by synthesizing an oligonucleotide on the basis of base sequences of portion in the cDNA of the present invention and screening the cDNA libraries using this oligonucleotide as a probe for colony or plaque hybridization according to a method known

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in the art. In addition, the cDNA fragments of the present invention can be prepared from an mRNA isolated from human cells by the RT-PCR method in which oligonucleotides which hybridize with both termini of the cDNA fragment of interest are synthesized, which oligonucleotides are then used as the primers.

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The cDNAs of the present invention are characterized in that they comprise any one of the base sequences represented by SEQ ID NOS: 11 to 20, 41 to 50, 71 to 80, 101 to 110 and 131 to 140 or the base sequences represented by SEQ ID NOS: 21 to 30, 51 to 60, 81 to 90, 111 to 120 and 141 to 150. Tables 1 and 2 summarizes the clone number (HP number), the cell from which the cDNA clone was obtained, the total number of bases of the cDNA, and the number of the amino acid residues of the encoded protein, for each of the cDNAs.

Table 1

SEQ	ID NO.		HP Cell	Cell	Number of	Number of amino acid
					bases	residues
1,	11,	21	HP03171	Thymus	2042	267
2,	12,	22	HP03424	Liver	1433	419
3,	13,	23	HP03444	Kidney	1917	415
4,	14,	24	HP03478	Umbilical cord blood	2258	380
5,	15,	25	HP03499	Kidney	1973	585
6,	16,	26	HP03500	kidney	1606	331
7,	17,	27	HP10691	Umbilical cord blood	2380	345
8,	18,	28	HP10703	Kidney	2017	89
9,	19,	29	HP10711	Kidney	1606	406
10,	20,	30	HP10712	Kidney	1695	192
31,	41,	51	HP03010	Kidney	1551	377
32,	42,	52	HP03576	Kidney	1713	81
33,	43,	53	HP03611	Kidney	1758	487
34,	44,	54	HP03612	Kidney	1550	375
35,	45,	55	HP10407	Stomach cancer	1485	350
36,	46,	56	HP10713	Kidney	2694	667
37,	47,	57	HP10714	Umbilical cord blood	3297	464
38,	48,	58	HP10716	Umbilical cord blood	2126	470
39,	49,	59	HP10717	Kidney	1781	243
40,	50,	60	HP10718	Umbilical cord blood	1788	270
61,	71,	81	HP03745	Kidney	1376	389
62,	72,	82	HP03747	Umbilical cord blood	2392	348
63,	73,	83	HP10719	Kidney	1416	261
64,	74,	84	HP10720	Kidney	1347	222
65,	75,	85	HP10721	Kidney	2284	183

Table 2

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Table						
SEQ	ID 1	NO	HP number	Cell	Number of bases	Number of amino acid residues
66,	76,	86	HP10725	Kidney	1737	262
67,	77,	87	HP10727	_	1556	168
68,	78,	88	HP10728	Umbilical cord blood	1855	243
69,	79,	89	HP10730	Umbilical cord blood		428
70,	80,	90	HP10742	Umbilical cord blood	1911	283
91,	101,	111	HP03800	Umbilical cord blood	1633	476
92,	102,	112	HP03831	Kidney	1095	226
93,	103,	113	HP03879	Kidney	1602	305
94,	104,	114	HP03880	Kidney	897	227
95,	105,	115	HP10704	Kidney	1866	441
96,	106,	116	HP10715	Umbilical cord blood	2198	265
97,	107,	117	HP10724	Umbilical cord blood	2180	208
98,	108,	118	HP10733	Umbilical cord blood	1527	400
99,	109,	119	HP10734	Umbilical cord blood	1905	192
100,	110,	120	HP10756	Kidney	998	260
121,	131,	141	HP03670	Umbilical cord blood	1622	337
122,	132,	142	HP03688	Umbilical cord blood	2475	236
123,	133,	143	HP03825	Kidney	1739	560
124,	134,	144	HP03877	Kidney	2005	406
125,	135,	145	HP10765	Umbilical cord blood	1558	453
126,	136,	146	HP10766	Kidney	1005	59
127,	137,	147	HP10770	Kidney	969	210
128,	138,	148	HP10772	Kidney	1241	165
129,	139,	149	HP10773	Kidney	1174	162
130,	140,	150	HP10776	Kidney	1012	221

The same clones as the cDNAs of the present invention can be easily obtained by screening the cDNA libraries constructed from the human cell lines or human

tissues utilized in the present invention using an oligonucleotide probe synthesized on the basis of the base sequence of the cDNA provided in any one of SEQ ID NOS: 11 to 30, 41 to 60, 71 to 90, 101 to 120 and 131 to 150.

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In general, the polymorphism due to the individual differences is frequently observed in human genes. Accordingly, any cDNA in which one or plural nucleotides are added, deleted and/or substituted with other nucleotides in SEQ ID NOS: 11 to 30, 41 to 60, 71 to 90, 101 to 120 and 131 to 150 shall come within the scope of the present invention.

Similarly, any protein in which one or plural amino acids are added, deleted and/or substituted with other amino acids resulting from the above-mentioned changes shall come within the scope of the present invention, as long as the protein possesses the activity of the protein having any one of the amino acid sequences represented by SEQ ID NOS: 1 to 10, 31 to 40, 61 to 70, 91 to 100 and 121 to 130.

The cDNAs of the present invention also include cDNA fragments (of 10 bp or more) containing any partial base sequence in the base sequences represented by SEQ ID NOS: 11 to 20, 41 to 50, 71 to 80, 101 to 110 and 131 to 140 or in the base sequences represented by SEQ ID NOS: 21 to 30, 51 to 60, 81 to 90, 111 to 120 and 141 to 150. Also, DNA fragments consisting of a sense strand and an anti-sense strand shall come within this scope. These DNA fragments can

be utilized as the probes for the genetic diagnosis.

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The antibody of the present invention can be obtained from a serum after immunizing an animal using the protein of the present invention as an antigen. A peptide that is chemically synthesized based on the amino acid sequence of the present invention and a protein expressed in eukaryotic or prokaryotic cells can be used as an antigen. Alternatively, an antibody can be prepared by introducing the above-mentioned expression vector for eukaryotic cells into the muscle or the skin of an animal by injection or by using a gene gun and then collecting a serum therefrom (JP-A 7-313187). Animals that can be used include a mouse, a rat, a rabbit, a goat, a chicken and the like. A monoclonal antibody directed to the protein of the present invention can be produced by fusing B cells collected from the spleen of the immunized animal with myelomas to generate hybridomas.

In addition to the activities and uses described above, the polynucleotides and proteins of the present invention may exhibit one or more of the uses or biological activities (including those associated with assays cited herein) identified below. Uses or activities described for proteins of the present invention may be provided by administration or use of such proteins or by administration or use of polynucleotides encoding such proteins (such as, for example, in gene therapies or vectors suitable for

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introduction of DNA).

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Research Uses and Utilities

polynucleotides provided bv the invention can be used by the research community for various purposes. The polynucleotides can be used to express recombinant protein for analysis, characterization therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either at a particular constitutively or stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to compare with endogenous DNA in patients to identify potential sequences disorders; as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR primers for genetic fingerprinting; as a probe to "subtract-out" known sequences in the process of discovering other novel polynucleotides; for selecting and making oligomers for attachment to a "gene chip" or other support, including for examination of expression patterns; to raise anti-protein antibodies using DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or potentially binds to another protein

(such as, for example, in a receptor-ligand interaction), the polynucleotide can also be used in interaction trap assays (such as, for example, that described in Gyuris et al., Cell '75:791-803 (1993)) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

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The proteins provided by the present invention can similarly be used in assay to determine biological activity, including in a panel of multiple proteins for highthroughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art. References disclosing such methods include without limitation "Molecular Cloning: A Laboratory Manual", 2d ed., Cold Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology: Guide to Molecular Cloning Techniques", Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

Nutritional Uses

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Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a nitrogen source and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

Activity

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A protein of the present invention may exhibit cytokine, cell proliferation (either inducing or inhibiting) or cell differentiation (either inducing or inhibiting) activity or may induce production of other cytokines in certain cell populations. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein of the present invention is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B9, B9/11, BaF3, MC9/G, M+ (preB M+), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7e and CMK.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for T-cell or thymocyte proliferation include without limitation those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Takai et al., J. Immunol. 137:3494-3500, 1986; Bertagnolli et al., J. Immunol.

145:1706-1712, 1990; Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Bertagnolli, et al., J. Immunol. 149:3778-3783, 1992; Bowman et al., J. Immunol. 152: 1756-1761, 1994.

- Assays for cytokine production and/or proliferation of spleen cells, lymph node cells or thymocytes include, without limitation, those described in: Polyclonal T cell stimulation, Kruisbeek, A.M. and Shevach, E.M. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 3.12.1-3.12.14, John Wiley and Sons, Toronto. 1994; and Measurement of mouse and human Interferon γ, Schreiber, R.D. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.8.1-6.8.8, John Wiley and Sons, Toronto. 1994.
- 15 Assays for proliferation and differentiation of hematopoietic and lymphopoietic cells include, without limitation, those described in: Measurement of Human and Murine Interleukin 2 and Interleukin 4, Bottomly, K., Davis, L.S. and Lipsky, P.E. In Current Protocols in Immunology.

 20 J.E.e.a. Coligan eds. Vol 1 pp. 6.3.1-6.3.12, John Wiley and Sons, Toronto. 1991; deVries et al., J. Exp. Med. 173:1205-1211, 1991; Moreau et al., Nature 336:690-692, 1988; Greenberger et al., Proc. Natl. Acad. Sci. U.S.A. 80:2931-2938, 1983; Measurement of mouse and human interleukin 6-Nordan, R. In Current Protocols in Immunology. J.E.e.a.

Coligan eds. Vol 1 pp. 6.6.1-6.6.5, John Wiley and Sons, Toronto. 1991; Smith et al., Proc. Natl. Acad. Sci. U.S.A. 83:1857-1861, 1986; Measurement of human Interleukin 11 - Bennett, F., Giannotti, J., Clark, S.C. and Turner, K. J. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.15.1 John Wiley and Sons, Toronto. 1991; Measurement of mouse and human Interleukin 9 - Ciarletta, A., Giannotti, J., Clark, S.C. and Turner, K.J. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.13.1, John Wiley and Sons, Toronto. 1991.

Assays for T-cell clone responses to antigens (which will identify, among others, proteins that affect APC-T cell interactions as well as direct T-cell effects by measuring proliferation and cytokine production) include, without limitation, those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function; Chapter 6, Cytokines and their cellular receptors; Chapter 7, Immunologic studies in Humans); Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

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A protein of the present invention may also exhibit immune stimulating or immune suppressing activity, including without limitation the activities for which assays are described herein. A protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined immunodeficiency (SCID)), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial or fungal infections, or may result from autoimmune disorders. More specifically, infectious diseases causes by viral, bacterial, fungal or other infection may be treatable using a protein of the present invention, including infections by HIV, hepatitis viruses, herpesviruses, mycobacteria, Leishmania spp., malaria spp. and various fungal infections such candidiasis. Of course, in this regard, a protein of the present invention may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Autoimmune disorders which may be treated using a protein of the present invention include, for example, connective tissue disease, multiple sclerosis, systemic lupus erythematosus, rheumatoid arthritis, autoimmune

pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, insulin dependent diabetes mellitis, myasthenia graft-versus-host disease and autoimmune gravis, inflammatory eye disease. Such a protein of the present invention may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems. conditions, in which immune suppression is (including, for example, organ transplantation), may also be treatable using a protein of the present invention.

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Using the proteins of the invention it may also be possible to immune responses, in a number of ways. Down regulation may be in the form of inhibiting or blocking an response already in progress or may involve immune induction of an immune response. preventing the functions of activated T cells may be inhibited by inducing specific suppressing T cell responses or by tolerance in T cells, or both. Immunosuppression of T cell responses is generally an active, non-antigen-specific, process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigenspecific and persists after exposure to the tolerizing agent has ceased. Operationally, tolerance can be demonstrated by

the lack of a T cell response upon reexposure to specific antigen in the absence of the tolerizing agent.

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Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions (such as , for example, B7)), e.g., preventing high level lymphokine synthesis by activated T cells, will situations of tissue, skin and organ be useful in transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue transplantation. in tissue transplants, rejection of Typically, transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2 activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen (e.g., B7-1, B7-3) or blocking antibody), prior to transplantation can lead to the binding of the molecule to the natural ligand(s) on the without transmitting the corresponding immune cells costimulatory signal. Blocking B lymphocyte antigen function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an immunosuppressant.

Moreover, the lack of costimulation may also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigen-blocking reagents may avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the function of a combination of B lymphocyte antigens.

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The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be 10 assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have 15 been used to examine the immunosuppressive effects of CTLA4Iq fusion proteins in vivo as described in Lenschow et al., Science 257:789-792 (1992) and Turka et al., Proc. Natl. Acad. Sci USA, 89:11102-11105 (1992). In addition, murine models of GVHD (see Paul ed., Fundamental Immunology, Raven 20 Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function in vivo on the development of that disease.

Blocking antigen function may also be therapeutically useful for treating autoimmune diseases.

Many autoimmune disorders are the result of inappropriate

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activation of T cells that are reactive against self tissue the production of cytokines which promote and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents T cells by disrupting which block costimulation of receptor: ligand interactions of B lymphocyte antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which may be involved in the disease process. Additionally, blocking antigen-specific tolerance induce reagents may autoreactive T cells which could lead to long-term relief from the disease. The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine encephalitis, systemic experimental autoimmune erythmatosis in MRL/lpr/lpr mice or NZB hybrid mice, murine autoimmune collagen arthritis, diabetes mellitus in NOD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy.

Upregulation of immune responses may be in the form of enhancing an existing immune response or eliciting an initial immune response. For example, enhancing an immune response through stimulating B lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the common cold, and encephalitis might be alleviated by the administration of stimulatory forms of B lymphocyte antigens systemically.

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alternatively, anti-viral immune responses may be enhanced in an infected patient by removing T cells from the patient, costimulating the T cells in vitro with viral antigen-pulsed APCs either expressing a peptide of the present invention or together with a stimulatory form of a soluble peptide of the present invention and reintroducing the in vitro activated T cells into the patient. Another method of enhancing anti-viral immune responses would be to isolate infected cells from a patient, transfect them with a nucleic acid encoding a protein of the present invention as described herein such that the cells express all or a portion of the protein on their surface, and reintroduce the transfected cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to, and thereby activate, T cells in vivo.

In another application, up regulation or enhancement of antigen function (preferably B lymphocyte

antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least one peptide of the present invention can be administered to a subject to overcome tumor-specific tolerance in the subject. If desired, the tumor cell can be transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected ex vivo with an expression vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection in vivo.

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The presence of the peptide of the present invention having the activity of a B lymphocyte antigen(s) on the surface of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack MHC class I or MHC class II molecules, or which fail to reexpress sufficient amounts of MHC class I or MHC class II molecules, can be transfected with nucleic acid encoding all or a portion of (e.g., a

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cytoplasmic-domain truncated portion) of an MHC class I lphachain protein and β , microglobulin protein or an MHC class II α chain protein and an MHC class II β chain protein to thereby express MHC class I or MHC class II proteins on the cell surface. Expression of the appropriate class I or class II MHC in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which blocks expression of an MHC class II associated invariant chain, protein, such as the can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject.

The activity of a protein of the invention may, among other means, be measured by the following methods:

20 Suitable assays for thymocyte or splenocyte cytotoxicity include, without limitation, those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19;

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Chapter 7, Immunologic studies in Humans); Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 5 1988; Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Bowmanet al., J. Virology 61:1992-1998; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 1994.

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Assays for T-cell-dependent immunoglobulin responses and isotype switching (which will identify, among others, proteins that modulate T-cell dependent antibody 15 responses and that affect Th1/Th2 profiles) include, without limitation, those described in: Maliszewski, J. Immunol. 144:3028-3033, 1990; and Assays for B cell function: In vitro antibody production, Mond, J.J. and Brunswick, M. In 20 Current Protocols in Immunology, J.E.e.a. Coligan eds. Vol 1 pp. 3.8.1-3.8.16, John Wiley and Sons, Toronto. 1994.

Mixed lymphocyte reaction (MLR) assays (which will identify, among others, proteins that generate predominantly Th1 and CTL responses) include, without limitation, those described in: Current Protocols in Immunology, Ed by J. E.

Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

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Dendritic cell-dependent assays (which will identify, among others, proteins expressed by dendritic cells that activate naive T-cells) include, without 10 limitation, those described in: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., Journal of Experimental Medicine 173:549-559, 1991; Macatonia et al., Journal of Immunology 154:5071-5079, 1995; Porgador et al., Journal of Experimental Medicine 182:255-260, 1995; Nair et al., 15 Journal of Virology 67:4062-4069, 1993; Huang et al., Science 264:961-965, 1994; Macatonia et al., Journal of Experimental Medicine 169:1255-1264, 1989; Bhardwaj et al., Journal of Clinical Investigation 94:797-807, 1994; and Inaba et al., Journal of Experimental Medicine 172:631-640, 20 1990.

Assays for lymphocyte survival/apoptosis (which will identify, among others, proteins that prevent apoptosis after superantigen induction and proteins that regulate lymphocyte homeostasis) include, without limitation, those

described in: Darzynkiewicz et al., Cytometry 13:795-808, 1992; Gorczyca et al., Leukemia 7:659-670, 1993; Gorczyca et al., Cancer Research 53:1945-1951, 1993; Itoh et al., Cell 66:233-243, 1991; Zacharchuk, Journal of Immunology 145:4037-4045, 1990; Zamai et al., Cytometry 14:891-897, 1993; Gorczyca et al., International Journal of Oncology 1:639-648, 1992.

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Assays for proteins that influence early steps of T-cell commitment and development include, without limitation, those described in: Antica et al., Blood 84:111-117, 1994; Fine et al., Cellular Immunology 155:111-122, 1994; Galy et al., Blood 85:2770-2778, 1995; Toki et al., Proc. Nat. Acad Sci. USA 88:7548-7551, 1991.

Hematopoiesis Regulating Activity

in regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or

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erythroid cells; in supporting the growth and proliferation of myeloid cells such as granulocytes monocytes/macrophages (i.e., traditional CSF activity) useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelo-suppression; in supporting growth and proliferation of megakaryocytes consequently of platelets thereby allowing prevention or various treatment of platelet disorders thrombocytopenia, and generally for use in place of or complementary to platelet transfusions; and/or in supporting the growth and proliferation of hematopoietic stem cells which are capable of maturing to any and all of the abovementioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantation, including, without limitation, aplastic anemia and paroxysmal nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in-vivo or conjunction with ex-vivo (i.e., in bone with peripheral progenitor transplantation or cell transplantation (homologous or heterologous)) as normal cells or genetically manipulated for gene therapy.

The activity of a protein of the invention may, among other means, be measured by the following methods:

25 Suitable assays for proliferation and

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differentiation of various hematopoietic lines are cited above.

Assays for embryonic stem cell differentiation (which will identify, among others, proteins that influence embryonic differentiation hematopoiesis) include, without limitation, those described in: Johansson et al. Cellular Biology 15:141-151, 1995; Keller et al., Molecular and Cellular Biology 13:473-486, 1993; McClanahan et al., Blood 81:2903-2915, 1993.

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10 Assays for stem cell survival and differentiation (which will identify, among others, proteins that regulate lympho-hematopoiesis) include, without limitation, those described in: Methylcellulose colony forming assays, Freshney, M.G. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 265-268, Wiley-Liss, Inc., New 15 York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; Primitive hematopoietic colony forming cells with high proliferative potential, McNiece, I.K. and Briddell, R.A. In Culture of Hematopoietic Cells. R.I. 20 Freshney, et al. eds. Vol pp. 23-39, Wiley-Liss, Inc., New York, NY. 1994; Neben et al., Experimental Hematology 22:353-359, 1994; Cobblestone area forming cell assay, Ploemacher, R.E. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 1-21, Wiley-Liss, Inc., New 25 York, NY. 1994; Long term bone marrow cultures in the presence of stromal cells, Spooncer, E., Dexter, M. and Allen, T. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 163-179, Wiley-Liss, Inc., New York, NY. 1994; Long term culture initiating cell assay, Sutherland, H.J. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 139-162, Wiley-Liss, Inc., New York, NY. 1994.

Tissue Growth Activity

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A protein of the present invention also may have utility in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.

A protein of the present invention, which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other animals. Such a preparation employing a protein of the invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. De novo bone formation induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth

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repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

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Another category of tissue regeneration activity that may be attributable to the protein of the present invention is tendon/ligament formation. A protein of the present invention, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue formation induced by composition of the present invention contributes to the repair of congenital, trauma induced, or other tendon or

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ligament defects of other origin, and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The compositions of the present invention may provide an environment to attract tendon or ligament-forming cells, stimulate growth of tendon- or ligament-forming cells, differentiation of progenitors of tendonligament-forming cells, or induce growth of tendon/ligament cells or progenitors ex vivo for return in vivo to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel The ligament defects. and other tendon or syndrome compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

The protein of the present invention may also be neural cells for proliferation of for useful regeneration of nerve and brain tissue, i.e. for the treatment of central and peripheral nervous system diseases and neuropathies, as well as mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein may be used in the treatment of diseases of the peripheral injuries, peripheral nerve such as nervous system, neuropathy and localized neuropathies, peripheral central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic

Shy-Drager syndrome. Further lateral sclerosis, and conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders, disorders, head trauma and such as spinal cord cerebrovascular diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

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Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds and the like.

It is expected that a protein of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to regenerate. A protein of the invention may also exhibit angiogenic activity.

A protein of the present invention may also be

useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokine damage.

A protein of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

The activity of a protein of the invention may, among other means, be measured by the following methods:

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Assays for tissue generation activity include, without limitation, those described in: International Patent Publication No. W095/16035 (bone, cartilage, tendon); International Patent Publication No. W095/05846 (nerve, neuronal); International Patent Publication No. W091/07491 (skin, endothelium).

Assays for wound healing activity include, without limitation, those described in: Winter, Epidermal Wound Healing, pps. 71-112 (Maibach, HI and Rovee, DT, eds.), Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978).

Activin/Inhibin Activity

A protein of the present invention may also exhibit activin- or inhibin-related activities. Inhibins are characterized by their ability to inhibit the release of

follicle stimulating hormone (FSH), while activins and are characterized by their ability to stimulate the release of follicle stimulating hormone (FSH). Thus, a protein of the present invention, alone or in heterodimers with a member of the inhibin α family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, as a homodimer or as a heterodimer with other protein subunits of the inhibin- β group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885. A protein of the invention may also be useful for advancement of the onset of fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

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The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for activin/inhibin activity include, without limitation, those described in: Vale et al., Endocrinology 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986; Vale et al., Nature 321:776-779, 1986; Mason et

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al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986.

Chemotactic/Chemokinetic Activity

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A protein of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, and/or endothelial epithelial eosinophils, Chemotactic and chemokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized example, attraction of lymphocytes, infections. For monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell

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chemotaxis.

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The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis) consist 5 of assays that measure the ability of a protein to induce the migration of cells across a membrane as well as the ability of a protein to induce the adhesion of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those 10 described in: Current Protocols in Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W.Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 6.12, Measurement of alpha and beta Chemokines 6.12.1-6.12.28; Taub et al. J. Clin. Invest. 15 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Muller et al Eur. J. Immunol. 25: 1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol. 153: 1762-1768, 1994.

20 Hemostatic and Thrombolytic Activity

A protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulation disorders (including hereditary disorders, such as hemophilias) or to enhance coagulation and other

hemostatic events in treating wounds resulting from trauma, surgery or other causes. A protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as, for example, infarction of cardiac and central nervous system vessels (e.g., stroke)).

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assay for hemostatic and thrombolytic activity include, without limitation, those described in: Linet et al., J. Clin. Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res. 45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79 (1991); Schaub, Prostaglandins 35:467-474, 1988.

15 Receptor/Ligand Activity

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A protein of the present invention may also demonstrate activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptor phosphatases and their ligands, receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and their ligands) and receptor/ligand pairs involved in antigen

presentation, antigen recognition and development of cellular and humoral immune responses). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein of the present invention (including, without limitation, fragments of receptors and ligands) may themselves be useful as inhibitors of receptor/ligand interactions.

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The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for receptor-ligand activity include without limitation those described in: Current Protocols in Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W.Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 7.28, Measurement of Cellular Adhesion under static conditions 7.28.1-7.28.22), Takai et al., Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al., J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp. Med. 169:149-160 1989; Stoltenborg et al., J. Immunol. Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995.

Anti-Inflammatory Activity

Proteins of the present invention may also exhibit anti-inflammatory activity. The anti-inflammatory activity

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may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cellcell interactions (such as, for example, cell adhesion), by inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can be used to treat inflammatory conditions including chronic or acute conditions), including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis, rejection, nephritis, hyperacute complement-mediated cytokine or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of cytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material.

Tumor Inhibition Activity

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In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly

(such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth.

Other Activities

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A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body example, breast (such as, for part size or shape augmentation or diminution, change in bone form or shape); effecting biorhythms or cardiac cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or elimination of dietary fat, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other

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nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent behaviors; providing analgesic effects or other reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulinlike activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antiqen in a vaccine composition to raise an immune response against such protein or another material or entity which is cross-reactive with such protein.

Examples

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The present invention is specifically illustrated
in more detail by the following Examples, but Examples are
not intended to restrict the present invention. The basic
procedures with regard to the recombinant DNA and the
enzymatic reactions were carried out according to the
literature ["Molecular Cloning. A Laboratory Manual", Cold

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Spring Harbor Laboratory, 1989]. Unless otherwise stated, restriction enzymes and various modifying enzymes to be used were those available from Takara Shuzo. The buffer compositions and the reaction conditions for each of the enzyme reactions were as described in the attached instructions. The cDNA synthesis was carried out according to the literature [Kato, S. et al., Gene 150: 243-250 (1994)].

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(1) Selection of cDNAs Encoding Proteins Having
Hydrophobic Domains

Human liver cDNA library (WO 98/21328) and human stomach cancer cDNA library (WO 98/21328), as well as the cDNA libraries constructed from human kidney mRNA (Clontech), human thymus mRNA (Clontech) and human umbilical cord blood mRNA were used as cDNA libraries.

Full-length cDNA clones were selected from the respective libraries and the whole base sequences thereof were determined to construct a homo-protein cDNA bank consisting of the full-length CDNA clones. The hydrophobicity/hydrophilicity profiles were determined for the proteins encoded by the full-length cDNA clones registered in the homo-protein cDNA bank by the Kyte-Doolittle method [Kyte, J. & Doolittle, R. F., J. Mol. Biol. 157: 105-132 (1982)] to examine the presence or absence of a hydrophobic domain. A clone that has a hydrophobic region

being assumed as a secretory signal or a transmembrane domain in the amino acid sequence of the encoded protein was selected as a clone candidate.

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(2) Protein Synthesis by In Vitro Translation

The plasmid vector bearing the cDNA of the present invention was used for in vitro transcription/translation with a T_NT rabbit reticulocyte lysate kit (Promega). In this case, [35S]methionine was added to label the expression product with a radioisotope. Each of the reactions was carried out according to the protocols attached to the kit. Two micrograms of the plasmid was subjected to the reaction at 30°C for 90 minutes in the reaction solution of a total volume of 25 μ l containing 12.5 μ l μ of T_NT rabbit reticulocyte lysate, 0.5 µl of a buffer solution (attached to the kit), 2 µl of an amino acid mixture (without methionine), 2 µl of [35S]methionine (Amersham) (0.37 MBq/µl), 0.5 µl of T7 RNA polymerase, and 20 U of RNasin. The experiment in the presence of a membrane system was carried out by adding 2.5 µl of a canine pancreas microsome fraction (Promega) to the reaction system. To 3 µl of the reaction solution was added 2 µl of the SDS sampling buffer (125 mM Tris-hydrochloride buffer, pH 6.8, 120 mM 2-mercaptoethanol, 2% SDS solution, 0.025% bromophenol blue and 20% glycerol) and the resulting mixture was heated at 95°C for 3 minutes and then subjected to SDS-polyacrylamide gel electrophoresis.

The molecular weight of the translation product was determined by carrying out the autoradiography.

(3) Expression in COS7

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Escherichia coli cells harboring the expression vector for the protein of the present invention were cultured at 37°C for 2 hours in 2 ml of the 2 x YT culture medium containing 100 μ g/ml of ampicillin, the helper phage M13K07 (50 μ 1) was added thereto, and the cells were then cultured at 37°C overnight. Single-stranded phage particles were obtained by polyethylene glycol precipitation from a supernatant separated by centrifugation. The particles were suspended in 100 μ l of 1 mM Tris-0.1 mM EDTA, pH 8 (TE).

The cultured cells derived from monkey kidney, COS7, were cultured at 37°C in the presence of 5% CO₂ in the Dulbecco's modified Eagle's medium (DMEM) containing 10% fetal calf serum. 1 x 10° COS7 cells were inoculated into a 6-well plate (Nunc, well diameter: 3 cm) and cultured at 37°C for 22 hours in the presence of 5% CO₂. After the medium was removed, the cell surface was washed with a phosphate buffer solution followed by DMEM containing 50 mM Trishydrochloride (pH 7.5) (TDMEM). A suspension containing 1 µl of the single-stranded phage suspension, 0.6 ml of the DMEM medium and 3 µl of TRANSFECTAM™ (IBF) was added to the cells and the cells were cultured at 37°C for 3 hours in the presence of 5% CO₂. After the sample solution was removed,

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the cell surface was washed with TDMEM, 2 ml per well of DMEM containing 10% fetal calf serum was added, and the cells were cultured at 37°C for 2 days in the presence of 5% CO₂. After the medium was exchanged for a medium containing [35S]cysteine or [35S]methionine, the cells were cultured for one hour. After the medium and the cells were separated each other by centrifugation, proteins in the medium fraction and the cell membrane fraction were subjected to SDS-PAGE.

(4) Preparation of Antibodies

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A plasmid vector containing the cDNA of the present invention was dissolved in a phosphate buffer solution (PBS: 145 mM NaCl, 2.68 mM KCl, 8.09 mM Na, HPO, 2 mM KH_2PO_4 , pH 7.2) to a concentration of 2 $\mu g/\mu l$. 25 μl each (a total of 50 µl) of the thus-prepared plasmid solution in PBS was injected into the right and left musculi quadriceps femoris of three mice (ICR line) using a 26 guage needle. After similar injections were repeated for one month at intervals of one week, blood was collected. The collected blood was stored at 4°C overnight to coagulate the blood, and then centrifuged at 8,000 x g for five minutes to obtain a supernatant. NaN, was added to the supernatant to a concentration of 0.01% and the mixture was then stored at The generation of an antibody was confirmed by 4°C. immunostaining of COS7 cells into which the corresponding vector had been introduced or by Western blotting using a

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cell lysate or a secreted product.

(5) Clone Examples

<HP03171> (SEQ ID NOS: 1, 11 and 21)

Determination of the whole base sequence of the 5 cDNA insert of clone HP03171 obtained from cDNA library of human thymus revealed the structure consisting of a 90-bp 5'-untranslated region, a 804-bp ORF, and a 1148-bp 3'untranslated region. The ORF encodes a protein consisting of 267 amino acid residues and there existed one putative 10 transmembrane domain. Figure 1 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 34 kDa that was somewhat larger than the molecular weight 15 of 30,234 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 38 kDa. In addition, there exists in the amino acid sequence of this protein one site at which N-glycosylation may occur (Asn-Thr-Thr at position 169).

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to chicken putative transmembrane protein E3-16 (Accession No. AAB70816). Table 3 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and chicken putative

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transmembrane protein E3-16 (GG). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 43.0% in the entire region.

Table 3

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HP RATRRINKRGAKNCNAIRHFENTFVVETLICGVV

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GG KEAMKGIQKREAVNCRKIRHFENRFAMETLICEQ

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AL036384) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03424> (SEQ ID NOS: 2, 12 and 22)

Determination of the whole base sequence of the cDNA insert of clone HP03424 obtained from cDNA library of human liver revealed the structure consisting of a 4-bp 5'-untranslated region, a 1260-bp ORF, and a 169-bp 3'-untranslated region. The ORF encodes a protein consisting of 419 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 2 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 50 kDa that was somewhat larger than the molecular weight

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of 46,375 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 54 kDa. In addition, there exist in the amino acid sequence of this protein six sites at which N-glycosylation may occur (Asn-Ala-Ser at position 29, Asn-Val-Thr at position 40, Asn-Cys-Thr at position 112, Asn-Lys-Ser at position 135, Asn-Ile-Ser at position 172 and Asn-Phe-Ser at position 189). Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from aspartic acid at position 28.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Drosophila melanogaster GOLIATH Table 4 shows protein (Accession No. 006003). comparison between amino acid sequences of the human protein of the present invention (HP) and Drosophila melanogaster GOLIATH protein (DM). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 40.8% in the intermediate region of 218 amino acid residues.

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Ta	ble 4
HP	MSCAGRAGPARLAALALLTCSLWPARADNASQEYYTALINVTVQEPGRGAPLTFRIDRGI
HP	YGLDSPKAEVRGQVLAPLPLHGVADHLGCDPQTRFFVPPNIKQWIALLQRGNCTFKEKIS
HP	RAAFHNAVAVVIYNNKSKEEPVTMTHPGTGDIIAVMITELRGKDILSYLEKNISVQMTI
DM	
HP	VGTRMPPKNFSRGSLVFVSISFIVLMIISSAWLIFYFIQKIRYTNARDRNQRRLGDA * **. * .******* *. *****. **** .*. *.**. ***
DM	EGRRGVRTISSLNRTSVLFVSISFIVDDILCWLIFYYIQRFRYMQAKDQQSRNLCSV
НР	KKAISKLTTRTVKKGDKETDPDFDHCAVCIESYKQNDVVRILPCKHVFHKSCVDPWLSE
DM	KKAIMKIPTKTGKFSD-EKDLDSDCCAICIEAYKPTDTIRILPCKHEFHKNCIDPWLIE
HP	CTCPMCKLNILKALGIVPNLPCTDNVAFDMERLTRTQAVNRRSALGDLAGDNSLGLEPL
DM	RTCPMCKLDVLKFYGYVVGDQIYQTPS—PQHTAPIASIEEVPVIVVAVPHGPQPLQPL
HP	TSGISPLPQDGELTPRTGEINIAVTKEWFIIASFGLLSALTLCYMIIRATASLNANEVE

DM ASNMSSFAPSHYFQSSRSPSSSVQQQLAPLTYQPHPQQAASERGRRNSAPATMPHAITAS

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HP F

DM HQVTDV

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA082118) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03444> (SEQ ID NOS: 3, 13 and 23)

Determination of the whole base sequence of the cDNA insert of clone HP03444 obtained from cDNA library of human kidney revealed the structure consisting of a 209-bp 5'-untranslated region, a 1248-bp ORF, and a 460-bp 3'-untranslated region. The ORF encodes a protein consisting of 415 amino acid residues and there existed a putative secretory signal at the N-terminus. Figure 3 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 43 kDa that was somewhat smaller than the molecular

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weight of 45,691 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 42 kDa. Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from glutamine at position 24.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human type I procollagen C-proteinase enhancer protein (Accession No. BAA23281). Table 5 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human type I procollagen C-proteinase enhancer protein (CP). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 43.6% in the entire region.

20 Table 5

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HP MRGANAWAPLCLLLAAATQLSRQQSPERPVFTCGGILTGESGFIGSEGFPGVYP

* **. * * **** *** . . *****. . **

CP MLPAATASLLGPLLTACALLPFA-Q-GQTPNYTRPVFLCGGDVKGESGYVASEGFPNLYP

	HP	PNSKCTWKITVPEGKVVVLNFRFIDLESDNLCRYDFVDVYNGH-ANGQRIGRFCGTFRPG
		. *. *. **** . * *. ** . *** *
	СР	PNKECIWTITVPEGQTVSLSFRVFDLELHPACRYDALEVFAGSGTSGQRLGRFCGTFRPA
5	НР	ALVSSGNKMMVQMISDANTAGNGFMAMFSAAEPNERGDQYCGGLLDRPSGSFKTPNWPDR
		. ** ** * * *
	CP	PLVAPGNQVTLRMTTDEGTGGRGFLLWYSGRATSGTEHQFCGGRLEKAQGTLTTPNWPES
	HР	DYPAGVTCVWHIVAPKNQLIELKFEKFDVERDNYCRYDYVAVFNGGEVNDARRIGKYCGD
10		***. *. * ***. ** . *. *. *. *. *. *. *.
	СР	DYPPGISCSWHIIAPPDQVIALTFEKFDLEPDTYCRYDSVSVFNGAVSDDSRRLGKFCGD
	HP	SPPAPIVSERNELLIQFLSDLSLTADGFIGHYIFRPKKLPTTTE
		. *. * ** **** **. ***** . * *
15	CP	AVPGSISSEGNELLVQFVSDLSVTADGFSASYKTLPRGTAKEGQGPGPKRGTEPKVKLPP
	НР	QPVTTTFPVTTGLKTTVALCQQKCRRTGTLEGNYCSSDFVLAGTVITTITRDG-SLHATV
20	CP	KSQPPEKTEESPSAPDAPTCPKQCRRTGTLQSNFCASSLVVTATVKSMVREPGEGLAVTV
·	HР	SIINIYKEGNLAIQQAGKNMSARLTVVCKQCPLLRRGLNYIIMGQVGEDGRGKIM-PNSF
		*. *. **. *. * * * * * * * * * * * * * * *
	CP	SLIGAYKTGGLDLPSPPTGASLKFYVPCKQCPPMKKGVSYLLMGQV-EENRGPVLPPESF

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CP VVLHRPNQDQILTNLSKRKCPSQPVRAAASQD

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. D78874) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03478> (SEQ ID NOS: 4, 14 and 24)

Determination of the whole base sequence of the cDNA insert of clone HP03478 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 224-bp 5'-untranslated region, a 1143-bp ORF, and a 891-bp 3'-untranslated region. The ORF encodes a protein consisting of 380 amino acid residues and there existed five putative transmembrane domains. Figure 4 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the

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protein was similar to Halocynthia roretzi HrPET-1 protein (Accession No. BAA81907). Table 6 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and Halocynthia roretzi HrPET-1 protein (HR). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 36.8% in the entire region.

Table 6

HP MLQTLYDYFWWERLWLPVNLTWADLEDRDGRVYAKASDLYITLPLALLFLIVRYFFEL

.*.**..** *..*** ********** ****...** * ...** * ...**

HR MDLLMDLYHWFWNEKFWLPQNLTWEDLKRTEEKQFGETRDLWLTFPLCITVLCIRFSVEK

HP YVATPLAALLNIKEKTRLRAPPNATLEHFYLTSGKQPKQVEVELLSRQSGLSGRQVERWF

.* **. **..* ... * .** ... * ... * ... * ... * * * **

20 HR GIARPLGKWLNLSERLHTPPRENIVLEKVYKTITRKPNYSQVEDLCKQTGWRKHEINVWF

HP RRRRNQDRPSLLKKFREASWRFTFYLIAFIAGMAVIVDKPWFYDMKKVWEGYPIQSTIPS

*... .**. *.**. ***. ***. * ..

HR RKKNLVGRPTTLTKFQETFWRFAFYLTSFFYGLYVMYDQECVWQTEKCFSNYPEDHVLSQ

	HP Q-YWYYMIELSI	YWSLLFS1ASD	VKKKDFKEQ1	THHAVITIETS:	F2MF4WATI	CAGILIMA
ŝ	. *.**.***.	k* *	***** * .	***. ****	*. **	**
	HR KIYYYYLIELAI	FYSATTLTQFFD	VKRKDFWEMF	IHHIVTIILLC	GSYTLNYTH	CMGAFILV
5	HP LHDSSDYLLES	AKMFNYAGWKNT	CNNIFIVFAI	VFIITRLVILP	FWILHCTL	/YPLELYP
	.***.** *	*** . **	* ** *. *	******	. **	*. *
	HR VHDSADFYIEF	AKMGKYANNSLV	TNVGFISFTI	SFFLSRLVILP	LWIVPSIW	YGIYTYN
	HP AFFGYYFFNSM	MGVLQLLHIFWA	YLILRMAHKF	ITGKLVEDERS	DREETESSI	EGEEAAAG
10	*	. *****. *	* *.	**	**.	k. *. * .
	HR CAMA-WLFCAL	L-ILQLLHFYWF	SHIVKAAYAS	ILVGVIERDTR	SESEDSSAI	EDETAKYS
	HP GGAKSRPLANG	HPILNNNHRKND			•	
	*.					
15	HR VGSGDYTESNG	IHKRVVTAR				

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T27334) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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Determination of the whole base sequence of the cDNA insert of clone HP03499 obtained from cDNA library of human kidney revealed the structure consisting of a 129-bp 5'-untranslated region, a 1758-bp ORF, and a 86-bp 3'untranslated region. The ORF encodes a protein consisting of 585 amino acid residues and there existed one putative transmembrane domain at the N-terminus. Figure 5 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. translation resulted in formation of a translation product of 63 kDa that was almost identical with the molecular weight of 63,987 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 82 kDa. In addition, there exist in the amino acid sequence of this protein five sites at which N-glycosylation may occur (Asn-Ile-Thr at position 89, Asn-Glu-Thr at position 106, Asn-Ala-Thr at position 189, Asn-Arg-Thr at position 220 and Asn-Ala-Thr at position 315).

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The search of the protein database using the amino 20 acid sequence of the present protein revealed that the protein was similar to Chinese hamster hypothetical protein 2BE2121 (Accession No. A30227). Table 7 shows comparison between amino acid sequences of the human protein the present invention (HP) and Chinese 25 hypothetical protein 2BE2121 (CH). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 44.8% in the entire region.

Table 7

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HP MVCREQLSKNQVKWVFAGITCVSVVVIAAIVLAITLRRPGCELEACSPDADMLDYLLSLG 10 ..***.*. . CH SWSENILDYFLRNS HP QISRRDALEVTWYHAANSKKAMTAALNSNITVLEADVNVEGLGTANETGVPIMAHPPTIY CH QITTEDGAEIIWYHAANHKSQMQEALRSAAHMIEADVLLPS--DGSEHGQPIMAHPPEMN 15 HP SDNTLEQWLDAVLGSSQKGIKLDFKNIKAVGPSLDLLRQLTEEGKVRRPIWINADILKGP CH SDNTLQEWLAEVM-KSNKGIKLDFKSLAAARASMLFLDNVKQH--LQCPVWMNADVLPGP 20 HP NMLISTEVNATQFLALVQEKYPKATLSPGWTTFYMSTSPNRTYTQAMVEKMHELVGGVPQ CH NG-SSKVVDAKAFLDTVTSFFPDVTFSLGWTTGWHPEKVNEGYSWTMVKEMDYICSGLTQ

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.*****...**... ***...*. ... ***.*. **...*. ... *****.** ... **...**

CH PVTFPVRAALVRQSCSQLLWLLKKSNRYSLTVWTGKDDSYPTEDLLYIRDYFNKTQVFYD

HP IFEPLLSQFKQLALNATRKPMYYTGGSLIPLLQLPGDDGLNVEWLVPDVQGSGKTATMTL

5 *.** .***

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CH ILEPQSHEFKQAIGI

Furthermore, the search of the GenBank using the

base sequences of the present cDNA has revealed the

registration of sequences that shared a homology of 90% or

more (for example, Accession No. R92398) among ESTs. However,

since they are partial sequences, it can not be judged

whether or not they encode the same protein as the protein

of the present invention.

<HP03500> (SEQ ID NOS: 6, 16 and 26)

Determination of the whole base sequence of the cDNA insert of clone HP03500 obtained from cDNA library of human kidney revealed the structure consisting of a 134-bp 5'-untranslated region, a 996-bp ORF, and a 476-bp 3'-untranslated region. The ORF encodes a protein consisting of 331 amino acid residues and there existed one putative transmembrane domain at the N-terminus. Figure 6 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro

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translation resulted in formation of a translation product of 38 kDa that was almost identical with the molecular weight of 37,694 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the amino acid sequence of the protein matched with that of human hypothetical protein (Accession No. AAC05803) in which a region of 62 amino acid residues from glycine at position 88 to lysine at position 149 was deleted.

of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA340631) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10691> (SEQ ID NOS: 7, 17 and 27)

Determination of the whole base sequence of the cDNA insert of clone HP10691 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 246-bp 5'-untranslated region, a 1038-bp ORF, and a 1096-bp 3'-untranslated region. The ORF encodes a protein consisting of 345 amino acid residues and there existed at least two putative transmembrane domains. Figure 7 depicts the hydrophobicity/hydrophilicity profile, obtained by the

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Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human BB1 protein (Accession No. AAB37433). Table 8 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human BB1 protein (BB). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The C-terminal region of 215 amino acid residues of the present protein shared a homology of 81.9% with the N-terminal region of human BB1 protein.

Table 8

HP MSPEEWTYLVVLLISIPIGFLFKKAGPGLKRWGAAAVGLGLTLFTCGPHTLHSLVTILGT

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HP WALIQAQPCSCHALALAWTFSYLLFFRALSLLGLPTPTPFTNAVQLLLTLKLVSLASEVQ

MASGFSKGPTLGLLRRALPDGDT-QLQLLLRGNHDRPVLPLPHLPGLAGAA

BB

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10 HP NIDCYSTDFCVRVRDGMRYWNMTVQWWLAQYIYKSAPARSYVLRL

BB NIDCYSTDFCVRVRDGMRYWNMTVQWWLAQYIYKSAPARSYVLRTAWTMLLSAYWHGLHP

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. W48653) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10703> (SEQ ID NOS: 8, 18 and 28)

Determination of the whole base sequence of the CDNA insert of clone HP10703 obtained from cDNA library of human kidney revealed the structure consisting of a 359-bp

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5'-untranslated region, a 270-bp ORF, and a 1388-bp 3'-untranslated region. The ORF encodes a protein consisting of 89 amino acid residues and there existed one putative transmembrane domain. Figure 8 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 18 kDa that was larger than the molecular weight of 10,469 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T08343) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10711> (SEQ ID NOS: 9, 19 and 29)

Determination of the whole base sequence of the cDNA insert of clone HP10711 obtained from cDNA library of human kidney revealed the structure consisting of a 29-bp 5'-untranslated region, a 1221-bp ORF, and a 356-bp 3'-untranslated region. The ORF encodes a protein consisting of 406 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain at the N-terminus. Figure 9 depicts the

hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In translation resulted in formation of a translation product of 44 kDa that was almost identical with the molecular weight of 43,836 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 58 kDa. In addition, there exist in the amino acid sequence of this protein seven sites at which N-glycosylation may occur (Asn-Ser-Thr at position 65, Asn-Trp-Ser at position 95, Asn-Val-Ser at position 134, Asn-Ile-Thr at position 159, Asn-Gly-Ser at position 187, Asn-Arg-Ser at position 230 and Asn-Leu-Thr at position 333). Application of the (-3,-1)rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from glutamic acid at position 36.

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The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to mouse kidney predominant protein (Accession No. BAA92527). Table 9 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and mouse kidney predominant protein (MM). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The

both proteins shared a homology of 79.9% in the entire region.

Table 9

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA362394) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10712> (SEQ ID NOS: 10, 20 and 30)

Determination of the whole base sequence of the cDNA insert of clone HP10712 obtained from cDNA library of human kidney revealed the structure consisting of a 52-bp 5'-untranslated region, a 579-bp ORF, and a 1064-bp 3'-untranslated region. The ORF encodes a protein consisting of 192 amino acid residues and there existed four putative transmembrane domains. Figure 10 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-

Doolittle method, of the present protein. translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to mouse calcium channel gamma 5 subunit (Accession No. CAB86387). Table 10 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and mouse calcium channel gamma 5 subunit (MM). Therein, the marks of -, *, and . represent a 10 gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 75.0% in the entire region. 15

Table 10

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HS MTAVGVQAQRPLGQRQPRRSFFESFIRTLIITCVALAVVLSSVSICDGHWLLAEDRLFGL MM MTAIGAQAHKLLGLKRPHRSFFESFIRTLIIVCTALAVVLSSVSICDGHWLLVEDHLFGL

HS WHFCTTTNQSVPICFRDLGQAHVPGLAVGMGLVRSVGALAVVAAIFGLEFLMVSQLCEDK MM WYFCTIGNHSEPHCLRDLSQAHMPGLAVGMGLARSVAAMAVVAAIFGLEMLIVSQVCEDV

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HS HSQCKWVMGSILLLVSFVLSSGGLLGFVILLRNQVTLIGFTLMFWCEFTASFLLFLNAIS

MM RSRRKWAIGSYLLLVAFILSSGGLLTFIILLKNQINLLGFTLMFWCEFTASFLFFLNAAS

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HS GLHINSITHPWE

*****. *. **.

MM GLHINSLTQPWDPPAGTLAYRKRGYDGTSLI

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA910339) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03010> (SEQ ID NOS: 31, 41 and 51)

Determination of the whole base sequence of the cDNA insert of clone HP03010 obtained from cDNA library of human kidney revealed the structure consisting of a 97-bp 5'-untranslated region, a 1134-bp ORF, and a 320-bp 3'-untranslated region. The ORF encodes a protein consisting of 377 amino acid residues and there existed at least eight putative transmembrane domains. Figure 11 depicts the

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hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 42 kDa that was almost identical with the molecular weight of 41,462 predicted from the ORF as well as a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Arabidopsis thaliana hypothetical protein (Accession No. AAC34490). Table 11 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and Arabidopsis thaliana hypothetical protein (AT). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 42.0% in the entire region other than the N-terminal region.

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Table 11

 $HP\ MDSALSDPHNGSAEAGGPTNSTTRPPSTPEGIALAYGSLLLMALLPIFFGALRSVRCARG$

* * *.

	HP	KNASDMPETITSRDAARFPIIASCTLLGLYLFFKIFSQEYINLLLSMYFFVLGILALSHT
		** * *** * **.*. **. * * . * . * . ******
	AT	VKDTPPTETMSKEHAMRFPLVGSAMLLSLFLLFKFLSKDLVNAVLTAYFFVLGIVALSAT
5		·
	HP	${\tt ISPFMNKFFPASFPNRQYQLLFTQGSGENKEEIINYEFDTKDLVCLGLSSIVGVWYLLRK}$
		. * *
	AT	LLPAIRRFLPNPWNDNLIVWRFPYFKSLEVEFTKSQVVAGIPGTFFCAWYAWKK
10	НР	HWIANNLFGLAFSLNGVELLHLNNVSTGCILLGGLFIYDVFWVFGTNVMVTVAKSFEAPI
		. * **. * * * * * ** **
	AT	${\tt HWLANNILGLSFCIQGIEMLSLGSFKTGAILLAGLFFYDIFWVFFTPVMVSVAKSFDAPI}$
	НР	${\tt KLVFPQDLLEKGLEANNFAMLGLGDVVIPGIFIALLLRFDISLKKNTHTYFYTSFAAYIF}$
15		**. **
	AT	${\tt KLLFPTGDALRPYSMLGLGDIVIPGIFVALALRFDVSRRRQPQ-YFTSAFIGYAV}$
	ΗР	${\tt GLGLTIFIMHIFKHAQPALLYLVPACIGFPVLVALAKGEVTEMFSYEESNPKDPAAVTES}$
		*. *** .*. *. ******* ***
20	AT	${\tt GVILTIVVMNWFQAAQPALLYIVPAVIGFLASHCIWNGDIKPLLAFDESKTEE-ATTDES}$
	HP	KEGTEASASKGLEKKEK
		**
	AT	KTSEEVNKAHDE

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA380429) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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<HP03576> (SEQ ID NOS: 32, 42 and 52)

10 Determination of the whole base sequence of the cDNA insert of clone HP03576 obtained from cDNA library of human kidney revealed the structure consisting of a 88-bp 5'-untranslated region, a 246-bp ORF, and a 1379-bp 3'untranslated region. The ORF encodes a protein consisting of 15 81 amino acid residues and there existed two putative transmembrane domains. Figure 12 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In translation resulted in formation of a translation product 20 of 20 kDa that was larger than the molecular weight of 9,178 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human vacuolar proton ATPase 9 kDa (Accession No. NP_003936). Table 12 shows the comparison

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between amino acid sequences of the human protein of the present invention (HP) and human vacuolar proton ATPase 9 kDa (VP). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 71.2% in the entire region.

10 Table 12

HP MTAHSFALPVIIFTTFWGLVGIAGPWFVPKGPNRGVIITMLVATAVCCYLFWLIAILAQL

VP MAYHGLTVPLIVMSVFWGFVGFLVPWFIPKGPNRGVIITMLVTCSVCCYLFWLIAILAQL

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HP NPLFGPQLKNETIWYVRFLWE

VP NPLFGPQLKNETIWYLKYHWP

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. W22566) among ESTs. However, since they are partial sequences, it can not be judged

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whether or not they encode the same protein as the protein of the present invention.

<HP03611> (SEQ ID NOS: 33, 43 and 53)

Determination of the whole base sequence of the cDNA insert of clone HP03611 obtained from cDNA library of human kidney revealed the structure consisting of a 189-bp 5'-untranslated region, a 1464-bp ORF, and a 105-bp 3'untranslated region. The ORF encodes a protein consisting of 487 amino acid residues and there existed eleven putative depicts the Figure 13 transmembrane domains. hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human cystine/glutamate transporter (Accession No. BAA82628). Table 13 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human cystine/glutamate transporter (CG). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology

of 43.8% in the entire region other than the N-terminal region.

Table 13 5 HP MGDTGLRKRREDEKSIQSQEPKTTSLQKELGLISGISIIVGTIIGS *.... *.... *... *.. *.. ****. CG MVRKPVVSTISKGGYLQGNVNGRLPSLGNKEPPGQEKVQLKRKVTLLRGVSIIIGTIIGA HP GIFVSPKSVLSNTEAVGPCLIIWAACGVLATLGALCFAELGTMITKSGGEYPYLMEAYGP 10 CG GIFISPKGYLQNTGSVGMSLTIWTVCGYLSLFGALSYAELGTTIKKSGGHYTYILEVFGP HP IPAYLFSWASLIVIKPTSFAIICLSFSEYVCAPFYVGCKPPQIVVKCLAAAAILFISTVN 15 CG LPAFVRVWVELLIIRPAATAVISLAFGRYILEPFFIQCEIPELAIKLITAVGITVVMVLN HP SLSVRLGSYVQNIFTAAKLVIVAIIIISGLVLLAQCNTKNFDNSFEGAQLSVGAISLAFY 20 CG SMSVSWSARIQIFLTFCKLTAILIIIVPGVMQLIKGQTQNFKDAFSGRDSSITRLPLAFY HP NGLWAYDGWNQLNYITEELRNPYRNLPLAIIIGIPLVTACYILMNVSYFTVMTATELLQS CG YGMYAYAGWFYLNFVTEEVENPEKTIPLAICISMAIVTIGYVLTNVAYFTTINAEELLLS

- HP QAVAVTFGDRVLYPASWIVPLFVAFSTIGAANGTCFTAGRLIYVAGREGHMLKVLSYISV
 .*****.*.*.* * **.***.*.*.*.*.**.****...** *
 CG NAVAVTFSERLLGNFSLAVPIFVALSCFGSMNGGVFAVSRLFYVASREGHLPEILSMIHV
- - HP ISKPITMHLQMLMEVVPPEEDPE
 .*. **. **...*** *.
- 15 CG MSEKITRTLQIILEVVPEEDKL

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. R07056) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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Determination of the whole base sequence of the cDNA insert of clone HP03612 obtained from cDNA library of human kidney revealed the structure consisting of a 153-bp 5'-untranslated region, a 1128-bp ORF, and a 269-bp 3'untranslated region. The ORF encodes a protein consisting of 375 amino acid residues and there existed seven putative domains. transmembrane Figure 14 depicts hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 39 kDa that was somewhat larger than the molecular weight of 37,930 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human monocarboxylate transporter (Accession No. AAC70919). Table 14 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human monocarboxylate transporter (MC). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 41.7% in the N-terminal region of 192 amino acid residues.

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Table 14

	HP	MTPQPAGPPDGGWGWVVAAAAFAINGLSYGLLRSLGLAFPDLAEHFDRSAQDTAW
		.*. *******.**. * * * *
5	MC	MPPMPSAPPVHPPPDGGWGWIVVGATFISIGFSYAFPKAVTVFFKEIQQIFHTTYSEIAW
	HP	ISALALAVQQAASPVGSALSTRWGARPVVMVGGVLASLGFVFSAFASGLLHLYLGLGLLA **. *** *. **. *. * *. **. *. *. *. *. *
	МС	ISSIMLAVMYAGGPVSSVLVNKYGSRPVVIAGGLLCCLGMVLASFSSSVVQLYLTMGFIT
10	HP	GFGWALVFAPALGTLSRYFSRRRVLAVGLALTGNGASSLLLAPALQLLLDTFGWRGALLL
	MC	* * * *** * * ** * * * *** * * *** GLGLAFNLQPALTIIGKYFYRKRPMANGLAMAGNPVFLSSLAPFNQYLFNTFGWKGSFLI
15	НР	LGAITLHLTPCGALLLPLVLPGDPPAPPRSPLAALGLSLFTRRAFSIFALGTALVGGGYF
	МС	** *. *.** LGSLLLNACVAGSLMRPLGPNQTTSKSKNKTGKTEDDSSPKKIKTKKSTWEKVNKYLDFS
	НР	VPYVHLAPRFRPGPGGIRSSAGGGRGCDGGCGRPAGLRVAGRPRLGAPPAAAGRIRGSDW
20	МС	LFKHRGFLIYLSGNVIMFLGFFAPIIFPAPYAKDQGIDEYSAAFLLSVMAFVDMFARPSV
	НР	AGAVGGGAGARGGRRRELGGSPAGRGCGLWAERGELRPAGFRCTPRAGGRRRCGAGHRAG
25	MC	GLIANSKYIRPRIQYFFSFAIMFNGVCHLLCPLAQDYTSLVLYAVFFGLGFGSVSSVLFE

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HP DDADEPRGAPGPSPVRLPKG

MC TLMDLYGAPRFSSAVGLYTIVECGPVLLGPPLAGKLVDLTGEYKYMYMSCGAIVVAASVW

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI742291) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10407> (SEQ ID NOS: 35, 45 and 55)

Determination of the whole base sequence of the cDNA insert of clone HP10407 obtained from cDNA library of human stomach cancer revealed the structure consisting of a 100-bp 5'-untranslated region, a 1053-bp ORF, and a 332-bp 3'-untranslated region. The ORF encodes a protein consisting of 350 amino acid residues and there existed at least four putative transmembrane domains. Figure 15 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein.

The search of the protein database using the amino acid sequence of the present protein revealed that the

protein was longer by 35 amino acid residues at the N-terminus than human hypothetical protein (Accession No. CAB43375).

Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of a clone beginning from the 117th base of the present cDNA (Accession No. AL050274).

<HP10713> (SEQ ID NOS: 36, 46 and 56)

Determination of the whole base sequence of the 10 cDNA insert of clone HP10713 obtained from cDNA library of human kidney revealed the structure consisting of a 79-bp 5'-untranslated region, a 2004-bp ORF, and a 611-bp 3'untranslated region. The ORF encodes a protein consisting of 667 amino acid residues and there existed nine putative 15 transmembrane domains. 16 Figure depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to mouse retinoic acid-responsive protein (Accession No. AAC16016). Table 15 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and mouse retinoic acid-

responsive protein (MM). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 74.1% in the entire region.

Table 15

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	нР	SKGLQSSYSEEYLKNLLCKKKLGSSYH-ISKHGFLSWARVCLKHCIYIPQPGFHLPLKLV
		*. ***. ***. ***. ***. * . * . **
	MM	SQGLQTSYSEKYLRTLLCPKKLDSCSHPASKRSLLSRAWAFSHHSIYTPQPGFRLPLKLV
5	HP	LSATLTGTAIYQVALLLLVGVVPTIQKVRAGVTTDVSYLLAGFGIVLSEDKQEVVELVKH
		·*************************************
	ММ	${\tt ISATLTGTATYQVALLLLVSVVPTVQKVRAGINTDVSYLLAGFGIVLSEDRQEVVELVKH}$
	HP	${\tt HLWALEVCYISALVLSCLLTFLVLMRSLVTHRTNLRALHRGAALDLSPLHRSPHPSRQAI}$
10		***. *. ******** ***. *. ***. ***. ******
	MM	${\tt HLWTVEACYISALVLSCASTFLLLIRSLRTHRANLQALHRGAALDLDPPLQSIHPSRQAI}$
	НР	${\tt FCWMSFSAYQTAFICLGLLVQQIIFFLGTTALAFLVLMPVLHGRNLLLFRSLESSWPFWL}$
		. ****. ***** ******. ******. ***** *. *
15	MM	${\tt VSWMSFCAYQTAFSCLGLLVQQVIFFLGTTSLAFLVFVPLLHGRNLLLLRSLESTWPFWL}$
	HP	${\tt TLALAVILQNMAAHWVFLETHDGHPQLTNRRVLYAATFLLFPLNVLVGAMVATWRVLLSA}$
		*. ******* *. **. *. **. *. *. *****. *.
	ММ	TVALAVILQNIAANWIFLRTHHGYPELTNRRMLCVATFLLFPINMLVGAIMAVWRVLISS
20		
	HP	LYNAIHLGQMDLSLLPPRAATLDPGYYTYRNFLKIEVSQSHPAMTAFCSLLLQAQSLLPR
		. ******* ***. ***. ***. **. **** ***. ***. * **
	MM	LYNTVHLGQMDLSLLPQRAASLDPGYHTYQNFLRIEASQSHPGVIAFCALLLHAPSPQPR
25	HP	TMAAPQDSLRPGEEDEGMQLLQTKDSMAKGARPGASRGRARWGLAYTLLHNPTLQVFRKT

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HP ALLGANGAQP

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MM ALTSAKANGTQP

of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI760170) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10714> (SEQ ID NOS: 37, 47 and 57)

Determination of the whole base sequence of the cDNA insert of clone HP10714 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 82-bp 5'-untranslated region, a 1395-bp ORF, and a 1820-bp 3'-untranslated region. The ORF encodes a protein consisting of 464 amino acid residues and there existed a putative secretory signal at the N-terminus. Figure 17 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In

vitro translation resulted in formation of a translation product of 49 kDa that was somewhat smaller than the molecular weight of 52,340 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 52 kDa. In addition, there exist in the amino acid sequence of this protein two sites at which N-glycosylation may occur (Asn-Ala-Thr at position 164 and Asn-Asp-Ser at position 320). Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from threonine at position 22.

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA861134) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10716> (SEQ ID NOS: 38, 48 and 58)

Determination of the whole base sequence of the cDNA insert of clone HP10716 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 60-bp 5'-untranslated region, a 1413-bp ORF, and a 653-bp 3'-untranslated region. The ORF encodes a protein consisting of 470 amino acid residues and there existed one

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putative transmembrane domain at the N-terminus. Figure 18 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 61 kDa that was larger than the molecular weight of 52,086 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human hypothetical protein CGI-90 (Accession No. AAD34085). Table 16 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human hypothetical protein CGI-90 (CG). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 32.4% in the entire region.

20 Table 16

HP MSRLGALGGARAGLGLLLGTAAGLGFLCLLYSQRWKRTQRHGRSQSLPNSLDYTQTSDPG

HP RHVMLLRAVPGGAGDASVLPSLPREGQEKVLDRLDFVLTSLVALRREVEELRSSLRGLAG

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	НР	EIVGEVRCHMEENQRVARRRRFPFVRERSDSTGSSSVYFTASSGATFTDAESEGGYTTAN
	CG	MALAARLWRLLPFRRGAAPGSRLPA
5	НР	AESDNERDSDKESEDGEDEVSCETVKMGRKDSLDLEEEAASGASSALEAGGSSGLEDVLP
	CG	GPSGSRGIAAPARFRGFEVMGNPGTFNRGLLLSALSYLGFETYQVISQAAVVHATAKVEE
	НР	LLQQADELHRGDEQGKREGFQLLLNNKLVYGSRQDFLWRLARAYSDMCELT-EEVSEKKS
10		.*.*** ** .* .*** . ******* .**
	CG	ILEQADYLYESGETEKLYQLLTQYKESEDAELLWRLARASRDVAQLSRTSEEEKKL
	НР	YALDGKEEAEAALEKGDESADCHLWYAVLCGQLAEHESIQRRIQSGFSFKEHVDKAIALQ * *. **** * * ***
15	CG	LVYEALEYAKRALEKNESSFASHKWYAICLSDVGDYEGIKAKIANAYIIKEHFEKAIELN
		PENPMAHFLLGRWCYQVSHLSWLEKKTATALLESPLSATVEDALQSFLKAEELQPGFSKA * *.* *** *
0.0	CG	PKDATSIHLMGIWCYTFAEMPWYQRRIAKMLFATPPSSTYEKALGYFHRAEQVDPNFYSK
20	НР	GRVYISKCYRELGKNSEARWWMKLALELPDVTKEDLAIQKDLEELEVILRD * * . * . * . * . * . * . * . *
	CG	NLLLLGKTYLKLHNKKLAAFWLMKAKDYPAHTEEDKQIQTEAAQLLTSFSEKN

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA852295) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10717> (SEQ ID NOS: 39, 49 and 59)

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Determination of the whole base sequence of the cDNA insert of clone HP10717 obtained from cDNA library of human kidney revealed the structure consisting of a 73-bp 5'-untranslated region, a 732-bp ORF, and a 976-bp 3'untranslated region. The ORF encodes a protein consisting of 243 amino acid residues and there existed two putative domains. Figure 19 depicts the transmembrane hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 36 kDa that was larger than the molecular weight of 26,270 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI478174) among ESTs. However, since they are partial sequences, it can not be judged whether or not they

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encode the same protein as the protein of the present invention.

<HP10718> (SEQ ID NOS: 40, 50 and 60)

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Determination of the whole base sequence of the cDNA insert of clone HP10718 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 86-bp 5'-untranslated region, a 813-bp ORF, and a 889-bp 3'-untranslated region. The ORF encodes a protein consisting of 270 amino acid residues and there existed three putative transmembrane domains. Figure 20 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 28 kDa that was smaller than the molecular weight of 31,116 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Caenorhabditis elegans hypothetical protein Y53C10A (Accession No. CAA22139). Table 17 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and Caenorhabditis elegans hypothetical protein Y53C10A (CE). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the

present invention, respectively. The both proteins shared a homology of 54.8% in the entire region other than the N-terminal region.

5 Table 17

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HP MAGAEDWPGQ

CE MTSSSAASSSTTTSSTMMPDENECLKKEEERFKSPDPAPTLDEEVDIDTLPSMLEDDPNG

HP QLELDEDEASCCRWGAQHAGARELAALYSPGKRLQEWCSVILCFSLIAHNLVHLLLLARW

CE NVVECDLGFKGPRWGPQHAGAKKLASMYSKEKRLQEKVSLFAAIFLFSIVFIN-LLLS-W

15 HP EDT--PLVILGVVAGALIADFLSGLVHWGADTWGSVELPIVGKAFIRPFREHHIDPTAIT

CE ESSIWVSVLVSAVLGIMTADFASGLVHWAADTFGSVE-TWFGRSFIRPFREHHVDPTAIT

HP RHDFIETNGDNCLVTLLPLLNMAYKFRTHSPEALEQ--LYPWECFVFCLIIFGTFTNQIH

CE RHDIVEVNGDNCMLCVGPLLWILYQQMTYQRDAITQWATFHW--YILLLGIYVALTNQIH

HP KWSHTYFGLPRWVTLLQDWHVILPRKHHRIHHVSPHETYFCITTGWLNYPLEKIGFWRRL

CE KWSHTYFGLPTWVVFLQKAHIILPRSHHKIHHISPHACYYCITTGWLNWPLEYIGFWRKM

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HP EDLIQGLTGEKPRADDMKWAQKIK

* ** . **. **. *** *..

CE EWVVTTVTGMQPREDDLKWATKLQ

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA176107) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention. In addition, the region from position 466 to position 778 of the cDNA of the present invention matched with the region from position 2 to position 314 of human ubiquitin-conjugating enzyme E2 variant 1 (Accession NO. NM_003349) although no match was observed in another region.

<HP03745> (SEQ ID NOS: 61, 71 and 81)

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Determination of the whole base sequence of the cDNA insert of clone HP03745 obtained from cDNA library of human kidney revealed the structure consisting of a 99-bp 5'-untranslated region, a 1170-bp ORF, and a 107-bp 3'-untranslated region. The ORF encodes a protein consisting of 389 amino acid residues and there existed at least nine

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putative transmembrane domains. Figure 21 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human solute carrier family 7 (Accession No. NP_003974). Table 18 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human solute carrier family 7 (SC). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 36.0% in the N-terminal region of 397 amino acid residues.

Table 18

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HP

MDRGEKIQLKRVFGYWWGTSFLLINIIG

SC MEAREPGRPTPTYHLVPNTSQSQVEEDVSSPPQRSSETMQLKKEISLLNGVSLVVGNMIG

	. ********** *. ** ***
	SC SGIFVSPKGVLVHT-ASYGMSLIVWAIGGLFSVVGALCYAELGTTITKSGASYAYILEAF
	HP GSTVAFLNLWTSLFLGSGVVAG-QALLLAEYSIQPFFPSCSVPKLPKKCLALAMLWIVGI
5	***. **
	SC GGFIAFIRLWVSLLVVEPTGQAIIAITFANYIIQPSFPSCDPPYLACRLLAAACICLLTF
	HP LTSRGVKEVTWLQIASSVLKVSILSFISLTGVVFLIRGKKENVERFQNAFDAELPDISHL
	** ** ** * * . *. * . * . *.** * *
10	SC VNCAYVKWGTRVQDTFTYAKVVALIAIIVMGLVKLCQGHSEHFQDAFEGSSWDMGNL
	HP IQAIFQGYFAYSGELKKPRTTIPKCIFTALPLVTVVYLLVNISYLTVLTPR
	* *.***
	SC SLALYSALFSYSGWDTLNFVTEEIKNPERNLPLAIGISMPIVTLIYILTNVAYYTVLNIS
15	
	HP EILSSDAVAITWADRAFPSLAWIMPFAISTSLFSNLLISIFKSSRPIYLASQEGQLPLLF
	******.*.*.**.* * ** *** ****.**.*
	SC DVLSSDAVAVTFADQTFGMFSWTIPIAVALSCFGGLNASIFASSRLFFVGSREGHLPDLI
20	HP NTLNSHS-SPFTAVLLLVTLGSLAIILTSLIDLINYIFFTGSLWSILLMIGILRRRYQEF
	. ****. * ** * ***
	SC SMIHIERFTPIPALLFNCTMALIYLIVEDVFQLINYFSFSYWFFVGLSVVGQLYLRWKEF
	HP NLSIPYKVKLDF
25	* * *

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SC KRPRPLKLSVFFPIVFCICSVFLVIVPLFTDTINSLIGIGIALSGVPFYFMGVYLPESRR

<HP03747> (SEQ ID NOS: 62, 72 and 82)

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Determination of the whole base sequence of the cDNA insert of clone HP03747 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 21-bp 5'-untranslated region, a 1047-bp ORF, and a 1324-bp 3'-untranslated region. The ORF encodes a protein consisting of 348 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain at the C-terminus. Figure 22 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 40 kDa that was almost identical with the molecular weight of 39,685 predicted from the ORF. Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from proline at position 39.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human endoplasmic reticulum glycoprotein (Accession No. NP_006807). Table 19 shows the comparison between amino acid sequences of the human protein

of the present invention (HP) and human endoplasmic reticulum glycoprotein (ER). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 54.1% in the entire region.

Table 19

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- ER MAAEGWIWRWGWGRRCLGRPGLLGPGPGPTTPLFLLLL-LGSVTADITDGNS-EHLK
- HP VHFKIHGQGKKNLHGDGLAIWYTKDRMQPGPVFGNMDKFVGLGVFVDTYPNEEKQQERVF

 ****. ** *********. *. ******. *. * **. * **. * ****

 ER VHFKVHGTGKKNLHGDGIALWYTRDRLVPGPVFGSKDNFHGLAIFLDTYPNDET-TERVF

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HP EWRDCIEVPGVRLPRGYYFGTSSITGDLSDNHDVISLKLFELTVERTPEEEKLHRDVFLP

.....****** *******. ********. ***...**

ER EWKNCIDITGVRLPTGYYFGASAGTGDLSDNHDIISMKLFQLMVEHTPDEESIDWTKIEP

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HP SVDNMKLP----EMTAPL--PPLSGLALFLIVFFSLVFSVFAIVIGIILYNKWQEQSRK

.. . **...**...*. * *....*..*.*

ER SVNFLKSPKDNVDDPTGNFRSGPLTGWRVFLLLLCALLGIVVCAVVGAVVFQKRQERN-K

10 HP RFY

ER RFY

15 Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA262924) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10719> (SEQ ID NOS: 63, 73 and 83)

Determination of the whole base sequence of the cDNA insert of clone HP10719 obtained from cDNA library of human kidney revealed the structure consisting of a 54-bp

5'-untranslated region, a 786-bp ORF, and a 576-bp 3'-untranslated region. The ORF encodes a protein consisting of 261 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 23 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 33 kDa that was larger than the molecular weight of 27,435 predicted from the ORF. Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from asparagine at position 19.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to mouse endomucin (Accession No. AAD05208). Table 20 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and mouse endomucin (MM). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 47.9% in the entire region.

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	HP	MELLQVTIL-FLLP-SIC-SSNSTGVL-EAANNSLVVTTTKPSITTPNTESLQKNVVTPT
		* ***. *. * ***. *. *
5	ММ	MRLLQATVLFFLLSNSLCHSEDGKDVQNDSIPTPAETSTTKASVTIPGIVSV-TNPNKPA
	НР	TGTTPKGTITNELLKMSLMSTATFLTSKDEGLKATTTDVRKNDSIISNVTVTSVTLPNAV
		. **. *. **
	ММ	DGTPPEGTTKSDVSQTSLVTTINSLTTPKHEVGTTTEGPLRNESSTMKITVPNTPTSNAN
10		
	НР	STLQSSKPKTETQSSIKTTEIPGSVLQPDASPSKTGTLTSIPVTIPENTSQSQVIGTEGG
		***. *. *. **
	ММ	STLPGSQNKITTQLLDALPKITATPSASLTTAHTMSLLQDTEDR
15	НР	KNASTSATSRSYSSIILPVVIALIVITLSVFVLVGLYRMCWKADPGTPENGNDQPQSDKE
		* *, *, , *, , ************************
	ММ	KIATTPSTTPSYSSIILPVVIALVVITLLVFTLVGLYRICWKRDPGTPENGNDQPQSDKE
	ΗР	SVKLLTVKTISHESGEHSAQGKTKN
20		**********
	ММ	SVKLLTVKTISHESGEHSAQGKTKN

The search of the GenBank using the base sequences

of the present cDNA has revealed the registration of

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sequences that shared a homology of 90% or more (for example, Accession No. AA486620) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10720> (SEQ ID NOS: 64, 74 and 84)

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Determination of the whole base sequence of the cDNA insert of clone HP10720 obtained from cDNA library of human kidney revealed the structure consisting of a 25-bp 5'-untranslated region, a 669-bp ORF, and a 653-bp 3'untranslated region. The ORF encodes a protein consisting of 222 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 24 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. translation resulted in formation of a translation product of 28 kDa that was somewhat larger than the molecular weight of 25,219 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 35 kDa. In addition, there exist in the amino acid sequence of this protein two sites at which N-glycosylation may occur (Asn-Val-Thr at position 76 and Asn-His-Thr at position 93). Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to

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expect that the mature protein starts from glutamic acid at position 15.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI792241) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

10 <HP10721> (SEQ ID NOS: 65, 75 and 85)

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Determination of the whole base sequence of the cDNA insert of clone HP10721 obtained from cDNA library of human kidney revealed the structure consisting of a 74-bp 5'-untranslated region, a 552-bp ORF, and a 1658-bp 3'-15 untranslated region. The ORF encodes a protein consisting of 183 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 25 depicts the hydrophobicity/hydrophilicity profile, obtained by the 20 Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 23 kDa that was somewhat larger than the molecular weight of 19,989 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 22 kDa. 25 Application of the (-3,-1) rule, a method for predicting the

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cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from glutamic acid at position 25.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. R27187) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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<HP10725> (SEQ ID NOS: 66, 76 and 86)

Determination of the whole base sequence of the cDNA insert of clone HP10725 obtained from cDNA library of human kidney revealed the structure consisting of a 235-bp 5'-untranslated region, a 789-bp ORF, and a 713-bp 3'untranslated region. The ORF encodes a protein consisting of 262 amino acid residues and there existed one putative domain. Figure 26 depicts transmembrane hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example,

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Accession No. AI127782) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10727> (SEQ ID NOS: 67, 77 and 87)

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Determination of the whole base sequence of the cDNA insert of clone HP10727 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 102-bp 5'-untranslated region, a 507-bp ORF, and a 947bp 3'-untranslated region. The ORF encodes a protein consisting of 168 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 27 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 24 kDa that was larger than the molecular weight of 17,822 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 23 kDa. Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from lysine at position 29.

The search of the GenBank using the base sequences

of the present cDNA has revealed the registration of

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sequences that shared a homology of 90% or more (for example, Accession No. R80316) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10728> (SEQ ID NOS: 68, 78 and 88)

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Determination of the whole base sequence of the cDNA insert of clone HP10728 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 221-bp 5'-untranslated region, a 732-bp ORF, and a 902bp 3'-untranslated region. The ORF encodes a protein consisting of 243 amino acid residues and there existed one putative transmembrane domain at the N-terminus. Figure 28 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 30 kDa that was larger than the molecular weight of 26,534 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. H23535) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present 25 invention.

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<HP10730> (SEQ ID NOS: 69, 79 and 89)

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Determination of the whole base sequence of the cDNA insert of clone HP10730 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 27-bp 5'-untranslated region, a 1287-bp ORF, and a 1216-bp 3'-untranslated region. The ORF encodes a protein consisting of 428 amino acid residues and there existed one putative transmembrane domain. Figure 29 depicts hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In translation resulted in formation of a translation product of 50 kDa that was somewhat larger than the molecular weight of 48,992 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. C19105) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10742> (SEQ ID NOS: 70, 80 and 90)

Determination of the whole base sequence of the cDNA insert of clone HP10742 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 231-bp 5'-untranslated region, a 852-bp ORF, and a 828-

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bp 3'-untranslated region. The ORF encodes a protein consisting of 283 amino acid residues and there existed two putative transmembrane domains. Figure 30 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 30 kDa that was smaller than the molecular weight of 31,629 predicted from the ORF.

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T35949) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03800> (SEQ ID NOS: 91, 101 and 111)

Determination of the whole base sequence of the cDNA insert of clone HP03800 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 67-bp 5'-untranslated region, a 1431-bp ORF, and a 135-bp 3'-untranslated region. The ORF encodes a protein consisting of 476 amino acid residues and there existed a putative secretory signal at the N-terminus. Figure 31 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In

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vitro translation resulted in formation of a translation product of 55 kDa that was almost identical with the molecular weight of 54,110 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 58 kDa. In addition, there exist in the amino acid sequence of this protein four sites at which N-glycosylation may occur (Asn-Lys-Thr at position 81, Asn-Met-Thr at position 132, Asn-Val-Thr at position 307 and Asn-Gln-Thr at position 346). Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from leucine at position 23.

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The search of the protein database using the amino acid sequence of the present protein revealed that the similar to mosquito vitellogenic protein was carboxypeptidase (Accession No. P42660). Table 21 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and mosquito vitellogenic carboxypeptidase (VC). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 44.5% in the entire region. In addition, the C-terminal portion beginning from alanine at position 182 matched with human probable carboxypeptidase (Accession No. AAC23787) except one amino acid residue.

Table 21 5 MVGAMWKVIVSLVLLMPGPCDGLFRSLYRSVSMPPK-GDSGQPLFLTPYIEAGKIQKG HP VC MVKFHLLVLIAFTCYTCSDATLWNPYKKLMRGSASPPRPGESGEPLFLTPLLQDGKIEEA HP RELSLYGPFPGLNMKSYAGFLTVNKTYNSNLFFWFFPAQIQPEDAPVVLWLQGGPGGSSM 10 VC RNKARVNHPMLSSVESYSGFMTVDAKHNSNLFFWYVPAKNNREQAPILVWLQGGPGASSL HP FGLFVEHGPYVVTSNMTLRDRDFPWTTTLSMLYIDNPVGTGFSFTDDTHGYAVNEDDVAR 15 VC FGMFEENGPFHIHRNKSVKQREYSWHQNHHMIYIDNPVGTGFSFTDSDEGYSTNEEHVGE HP DLYSALIQFFQIFPEYKNNDFYVTGESYAGKYVPAIAHLIHSLNPVREVKINLNGIAIGD VC NLMKFIQOFFVLFPNLLKHPFYISGESYGGKFVPAFGYAIH--NSQSQPKINLQGLAIGD 20 HP GYSDPESIIGGYAEFLYQIGLLDEKQKKYFQKQCHECIEHIRKQNWFEAFEILDKLLDGD

VC GYTDPLNQL-NYGEYLYELGLIDLNGRKKFDEDTAAAIACAERKDMNSANRLIQGLFDG-

- HP AEKKVWKIFKSDSEVAGYIRQAGDFHQVIIRGGGHILPYDQPLRAFDMINRFIYGKGWDP

 *... *.. *.** ... *... ***... ***

 VC ANRE---IYRVDGEIAGYKKRAGRLQEVLIRNAGHMVPRDQPKWAFDMITSFTHKNYL

HP YVG

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA095665) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03831> (SEQ ID NOS: 92, 102 and 112)

Determination of the whole base sequence of the con con insert of clone HP03831 obtained from con library of

human kidney revealed the structure consisting of a 191-bp 5'-untranslated region, a 681-bp ORF, and a 223-bp 3'-untranslated region. The ORF encodes a protein consisting of 226 amino acid residues and there existed four putative transmembrane domains. Figure 32 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human claudin-10 (Accession No. NP_008915). Table 22 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human claudin-10 (CD). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 76.2% in the entire region. The C-terminal region downstream from glycine at position 72 completely matched with that sequence.

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	HP	MSRAQ1WALVSGVGGFGALVAATTSNEWKV1TRASSV1TATWVYQGLWMNCAGNALGS
		* ** ***.****** *
	CD	MASTASEIIAFMVSISGWVLVSSTLPTDYWKVSTIDGTVITTATYWANLWKACVTDSTGV
5	HP	FHCRPHFTIFKVAGYIQACRGLMIAAVSLGFFGSIFALFGMKCTKVGGSDKAKAKIACLA
		. * **************************
	CD	SNCKDFPSMLALDGYIQACRGLMIAAVSLGFFGSIFALFGMKCTKVGGSDKAKAKIACLA
	HP	GIVFILSGLCSMTGCSLYANKITTEFFDPLFVEQKYELGAALFIGWAGASLCIIGGVIFC
10		*****************
	CD	GIVFILSGLCSMTGCSLYANKITTEFFDPLFVEQKYELGAALFIGWAGASLCIIGGVIFC
	HP	FSISDNNKTPRYTYNGATSVMSSRTKYHGGEDFKTTNPSKQFDKNAYV

15	CD	FSISDNNKTPRYTYNGATSVMSSRTKYHGGEDFKTTNPSKQFDKNAYV

Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. N41613) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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Determination of the whole base sequence of the cDNA insert of clone HP03879 obtained from cDNA library of human kidney revealed the structure consisting of a 33-bp 5'-untranslated region, a 918-bp ORF, and a 651-bp 3'-untranslated region. The ORF encodes a protein consisting of 305 amino acid residues and there existed one putative transmembrane domain at the N-terminus. Figure 33 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 34 kDa that was almost identical with the molecular weight of 34,073 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human NADH-cytochrome b5 reductase (Accession No. Y09501). Table 23 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human NADH-cytochrome b5 reductase (CT). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 63.5% in the entire region other than the N-terminal region.

Table 23

	HP	${\tt MGIQTSPVLLASLGVGLVTLLGLAVGSYLVRRSRRPQVTLLDPNEKYLLRLLDKTTVSHN}$
		* . ** * . ** . ** *** **.
5	CT	MGAQLSTLGHMVLFPVWFLYSLLMKLFQRS-TPAITLESPDIKYPLRLIDREIISHD
	HP	${\tt TKRFRFALPTAHHTLGLPVGKHIYLSTRIDGSLVIRPYTPVTSDEDQGYVDLVIKVYLKG}$
		*. ******* *. ******. *****. **. *
	CT	TRRFRFALPSPQHILGLPVGQHIYLSARIDGNLVVRPYTPISSDDDKGFVDLVIKVYFKD
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	HP	${\tt VHPKFPEGGKMSQYLDSLKVGDVVEFRGPSGLLTYTGKGHFNIQPNKKSPPEPRVAKKLG}$
		·***** ******* * · · · ** · ******* * *** * * · * · * · * · * * * * · · * · * · * · * · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · * · · · * · · · * ·
	СТ	THPKFPAGGKMSQYLESMQIGDTIEFRGPSGLLVYQGKGKFAIRPDKKSNPIIRTVKSVG
15	HP	MIAGGTGITPMLQLIRAILKVPEDPTQCFLLFANQTEKDIILREDLEELQARYPNRFKLW

	CT	MIAGGTGITPMLQVIRAIMKDPDDHTVCHLLFANQTEKDILLRPELEELRNKHSARFKLW
		. ·
	HP	FTLDHPPKDWAYSKGFVTADMIREHLPAPGDDVLVLLCGPPPMVQLACHPNLDKLGYSQK
20		. *** *. *. * *** ***. * ** ****. * * * **** *
	CT	YTLDRAPEAWDYGQGFVNEEMIRDHLPPPEEEPLVLMCGPPPMIQYACLPNLDHVGHPTE
	HP	MRFTY
		. *
25	CT	RCFVF

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. F06459) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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<HP03880> (SEQ ID NOS: 94, 104 and 114)

Determination of the whole base sequence of the cDNA insert of clone HP03880 obtained from cDNA library of human kidney revealed the structure consisting of a 98-bp 5'-untranslated region, a 684-bp ORF, and a 115-bp 3'-untranslated region. The ORF encodes a protein consisting of 227 amino acid residues and there existed a putative secretory signal at the N-terminus. Figure 34 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 28 kDa that was somewhat larger than the molecular weight of 25,717 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 27 kDa. Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to

expect that the mature protein starts from aspartic acid at position 23.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to rat phosphatidylethanolamine-binding protein (Accession No. P31044). Table 24 shows the comparison between amino acid sequences of the human protein of invention the present (HP) and phosphatidylethanolamine-binding protein (RN). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 37.6% in the region of 133 amino acid residues other than the N-terminal region.

Table 24

HP MGWTMRLVTAALLLGLMMVVTGDEDENSPCAHEALLDEDTLFCQGLEVFYPELGNIGCKV

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RN MAADISQWAGPLSLQEVDEPPQHALRVDYGGVTV

HP VPDCNNYRQKITSWMEPIVKFPGAVDGATYILVMVDPDAPSRAEPRQRFWRHWLVTDIKG

RN DELGKVLTPTQVMNRPSSISWDGLDPGKLYTLVLTDPDAPSRKDPKFREWHHFLVVNMKG

HP ADLKKGKIQGQELSAYQAPSPPAHSGFHRYQFFVYLQEGKV——ISLLP-KENKTRGSWK

.*..*. **.**..** ..** **..**

RN NDISSGTV——LSEYVGSGPPKDTGLHRYVWLVYEQEQPLNCDEPILSNKSGDNRGKFK

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HP MDRFLNRFHLGEPEASTQFMTQNYQDSPTLQAPRERASEPKHKNQAEIAAC

...* ...***.* *.* * *.*.

RN VESFRKKYHLGAPVAGTCFQAEWDDSVPKLHDQLAGK

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. H83784) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10704> (SEQ ID NOS: 95, 105 and 115)

Determination of the whole base sequence of the

CDNA insert of clone HP10704 obtained from cDNA library of
human kidney revealed the structure consisting of a 141-bp
5'-untranslated region, a 1326-bp ORF, and a 399-bp 3'untranslated region. The ORF encodes a protein consisting of
441 amino acid residues and there existed eight putative
transmembrane domains. Figure 35 depicts the

hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human unknown gene product (Accession No. AAC27544). Table 25 shows the comparison between amino acid sequences of the human protein of the present invention

(HP) and human unknown gene product (UP). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 39.1% in the entire region.

Table 25

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HP MAIHKALVMCLGLPLFLFPG-AWAQGHVPPGCSQGLNPLYYNLCDRSGAWGIVLE

* **... *** * .***. *

UN MFVASERKMRAHQVLTFLLLFVITSVASENASTSRGCGLDLLPQYVSLCDLDAIWGIVVE

	HP	TCASRRFLFGVLFAICFSCLAAHVFALNFLARKNHGPRGWVIFTVALLLTLVEVIINTEW
		.*. ****.****. ***** *. *. ** ** ** * **. **
	UN	ICSVRRFLWGVLFALCFSCLLSQAWRVRRLVRHGTGPAGWQLVGLALCLMLVQVIIAVEW
5		
	HP	LIITLVRGSGEGGPQGNSSAGWAVASPCAIANMDFVMALIYVMLLLLGAFLGAWPALCGR
		*** * .******* *.** * .***.
	UN	LVLTVLRDTRPACAYEPMDFVMALIYDMVLLVVTLGLALFTLCGK
10	HP	YKRWRKHGVFVLLTTATSVAIWVVWIVMYTYGN-KQHNSPTWDDPTLAIALAANAWAFVL
		.****.*.*. ** ***.*. ** *
	UN	FKRWKLNGAFLLITAFLSVLIWVAWMTMYLFGNVKLQQGDAWNDPTLAITLAASGWVFVI
		·
	ΗР	FYVIPEVSQVTKSSPEQSYQGDMYPTRGVGY-ETILKEQ-KGQSMFVENKAFSMDEPVAA
15		****. * *
	UN	FHAIPEI-HCTLLPALQENTPNYFDTSQPRMRETAFEEDVQLPRAYMENKAFSMDEHNAA
	НР	KRPVS-PYSGYNGQLLTSVYQPTEMALMHKVPSEGAYDIILPRATANSQVMGSANSTLRA
		* *
20	UN	LRTAGFPNGSLGKRPSGSLGKRPSAPFRSNVYQPTEMAVVLNGGTIPTAPPSHTGRHLW
	НР	EDMYSAQSHQAATPPKDGKNSQVFRNPYVWD

of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA346702) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10715> (SEQ ID NOS: 96, 106 and 116)

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Determination of the whole base sequence of the cDNA insert of clone HP10715 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 49-bp 5'-untranslated region, a 798-bp ORF, and a 1351-bp 3'-untranslated region. The ORF encodes a protein consisting of 265 amino acid residues and there existed two putative transmembrane domains. Figure 36 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 43 kDa that was larger than the molecular weight of 29,217 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI381750) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present

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<HP10724> (SEQ ID NOS: 97, 107 and 117)

Determination of the whole base sequence of the cDNA insert of clone HP10724 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 68-bp 5'-untranslated region, a 627-bp ORF, and a 1485-bp 3'-untranslated region. The ORF encodes a protein consisting of 208 amino acid residues and there existed one putative transmembrane domain at the N-terminus. Figure 37 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 24 kDa that was almost identical with the molecular weight of 23,850 predicted from the ORF.

of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T78035) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10733> (SEQ ID NOS: 98, 108 and 118)

Determination of the whole base sequence of the cDNA insert of clone HP10733 obtained from cDNA library of human umbilical cord blood revealed the structure consisting

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of a 102-bp 5'-untranslated region, a 1203-bp ORF, and a 222-bp 3'-untranslated region. The ORF encodes a protein consisting of 400 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 38 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 50 kDa that was larger than the molecular weight of 43,151 predicted from the ORF. In this case, the addition of a microsome led to the formation of a product of 54 kDa. In addition, there exist in the amino acid sequence of this protein four sites at which N-qlycosylation may occur (Asn-Leu-Thr at position 52, Asn-Ala-Ser at position 131, Asn-Ile-Thr at position 145 and Asn-Leu-Ser at position 343). Application of the (-3,-1) rule, a method for predicting the cleavage site of the secretory signal sequence, allows to expect that the mature protein starts from arginine at position 33.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Drosophila melanogaster GOLIATH protein (Accession No. Q06003). Table 26 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and Drosophila melanogaster

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GOLIATH protein (DM). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 35.0% in the entire region.

Table 26

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HP MAWRREASVGARGVLALALLALALCVPGARGRALEWFSAVVNIEYVDPQTNLTVWSVSE

HP SGRFGDSSPKEGAHGLVGVPWAPGGDLEGCAPDTRFFVPEPGGRGAAPWVALVARGGCTF

HP KDKVLVAARRNASAVVLYNEERYGNITLPMSHAGTGNIVVIMISYPKGREILEL-VQKGI

* *... **. **. ***

DM MQLEKMQIKGKTRNIAAVITYQNIGQDLSLTLDKGY

HP PVTMTIGVGTRHVQEF--ISGQSVVFVAIAFITMMIISLAWLIFYYIQRFLY-TGSQIGS

..* * * *... **. **. * ******** * ... *

20 DM NVTISIIEGRRGVRTISSLNRTSVLFVSISFI--VDDILCWLIFYYIQRFRYMQAKDQQS

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	HP	DPWLLDHRICPMCKLDVIKALGYWGEPGDVQEMPAPESPPGKDPAANLSLALPDDDG5DE
		****. ********* ** *.*
	DM	DPWLIEHRTCPMCKLDVLKFYGY-VVGDQIYQTPSPQHTAPIASIEEVPVIVVAVPHGPQ
5	ΗР	SSPPSASPAESEPQCDPSFKGDAGENTALLEAGRSDSRHGGPIS
		* * *
	DM	PLQPLQASNMSSFAPSHYFQSSRSPSSSVQQQLAPLTYQPHPQQAASERGRRNSAPATMP

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI286184) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10734> (SEQ ID NOS: 99, 109 and 119)

Determination of the whole base sequence of the cDNA insert of clone HP10734 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 124-bp 5'-untranslated region, a 579-bp ORF, and a 1202-bp 3'-untranslated region. The ORF encodes a protein consisting of 192 amino acid residues and there existed one putative transmembrane domain. Figure 39 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-

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Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human sodium channel \$2\$ subunit (Accession No. AAD47196). Table 27 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human sodium channel \$2\$ subunit (SC). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 26.3% in the N-terminal region of 152 amino acid residues.

Table 27

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HP MFCPLKLILLPVLLDYSLGLNDLNVS-PPELTVHVGDSALMGCVFQS--TEDK

20 ...*. *....* *...* *...* *...* *...* *...

SC MHRDAWLPRPAFSLTGLSLFFSLVPPGRSMEVTVPATLNVLNGSDARLPCTFNSCYTVNH

HP CIFKIDWTLSPGEHAKDE-YVLYYYSNLSVPIGRFQNRVHLMGDNLCNDGSLLLQDVQEA

*...** ...* *...* *...* *...**...**

25 SC KQFSLNWTYQECNNCSEEMFLQFRMKIINLKLERFQDRVEFSGNPSKYDVSVMLRNVQPE

134

HP DQGTYICEIRLKGESQVFKKAVVLHVLPEEPKELMVHVGGLIQMGCVFQSTEVKHVTKVE

SC DEGIYNCYIMNPPDRHRGHGKIHLQVLMEEPPERDFTVAVIVGASVGGFLAVVILVLMVV

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HP WIFSGRRAKVTRRKHHCVREGSG

SC KCVRRKKEQKLSTDDLKTEEEGKTDGEGNPDDGAK

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. C03216) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10756> (SEQ ID NOS: 100, 110 and 120)

Determination of the whole base sequence of the cDNA insert of clone HP10756 obtained from cDNA library of human kidney revealed the structure consisting of a 49-bp 5'-untranslated region, a 783-bp ORF, and a 166-bp 3'-untranslated region. The ORF encodes a protein consisting of 260 amino acid residues and there existed a putative secretory signal at the N-terminus. Figure 40 depicts the

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hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 27 kDa that was almost identical with the molecular weight of 27,356 predicted from the ORF.

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AW027769) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03670> (SEQ ID NOS: 121, 131 and 141)

Determination of the whole base sequence of the cDNA insert of clone HP03670 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 77-bp 5'-untranslated region, a 1014-bp ORF, and a 531-bp 3'-untranslated region. The ORF encodes a protein consisting of 337 amino acid residues and there existed at least seven putative transmembrane domains. Figure 41 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to human hypothetical protein KIAA0260

(Accession No. BAA13390). Table 28 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and human hypothetical protein KIAA0260 (KI). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 57.6% in the entire region other than the N-terminal region. In addition, the C-terminal region beginning from leucine at position 77 matched with human putative Sqv-7-like protein (Accession No. AJ005866) except one amino acid residue.

Table 28

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HP

MTAGGQAEAEGAGGEPG

- KI NSWSPLGAAAAGPRAARPRRQATAAAAAMAEVHRRQHARVKGEAPAKSSTLRDEEELGMA
- - HP NKIIHFPDFDKKIPVKLFPLPLLYVGNHISGLSSTSKLSLPMFTVLRKFTIPLTLLLETI

137

KI LRVVKFPDLDRNVPRKTFPLPLLYFGNQITGLFSTKKLNLPMFTVLRRFSILFTMFAEGV

- 5 KI LLKKTFSWGIKMTVFAMIIGAFVAASSDLAFDLEGYAFILINDVLTAANGAYVKQKLDSK
- 15 HP SSQLKPKPVGEENICLDLKS
 *..*.***.*.
 KI TEEQLSKQ-SEANNKLDIKGKGAV

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. R24922) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present

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<HP03688> (SEQ ID NOS: 122, 132 and 142)

Determination of the whole base sequence of the cDNA insert of clone HP03688 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 35-bp 5'-untranslated region, a 711-bp ORF, and a 1729-bp 3'-untranslated region. The ORF encodes a protein consisting of 236 amino acid residues and there existed five putative transmembrane domains. Figure 42 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Caenorhabditis elegans hypothetical protein W02D9 (Accession No. CAB03470). Table 29 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and Caenorhabditis elegans hypothetical protein W02D9 (CE). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 50.8% in the entire region other than the N-terminal

re	egion.
Tá	able 29
НЕ	MAEAEI
CE	MEILNLSSKFSLSDKPCQKFIFSLFSAVQNSRFKIISFPEIHQKPLPQEEMNSFGNASVI
НІ	P SPGDPGTASPRPLFAGLSDISISQDIPVEGEITIPMRSRIREFDSSTLNESVRNTIMRD
	** **. *. **.
CI	IDMLEQEMAAEQTANLSGNIAGMSAPKSSSNRRGPMQEVDLDAEFDTLEEPVWDTVKRD
H	KAVGKKFMHVLYPR-KSNTLLRDWDLWGPLILCVTLALMLQRDSADSEKDGGPQFAEVF
	·** ** ** _· * _· ··· ********* _· ** _· *
CI	E LTVGAKFTHVVLPHGDKQQLLRDWDLWGPLFICVGLALLLQHNGGTESAPQFTQVF
Н	P IVWFGAVTITLNSKLLGGNISFFQSLCVLGYCILPLTVAMLICRLVLLADPGPVNFMVR
	* * * * * * * * * * * * * * * * * *
C	E ITFFGSVIVTANIKLLGGNISFFQSLCVIGYCLLPPFVAAVLCSL-FLHGIAFPLR
Н	P FVVIVMFAWSIVASTAFLADSQPPNRRALAVYPVFLFYFVISWMILTFTPQ
	*.**. ** .****** *********
C	E LITSIGFVWSTYASMGFLAGCQPDKKRLLVIYPVFLFYFVVSWMIISHS

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Furthermore, the search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T51465) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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<HP03825> (SEQ ID NOS: 123, 133 and 143)

Determination of the whole base sequence of the 10 cDNA insert of clone HP03825 obtained from cDNA library of human kidney revealed the structure consisting of a 20-bp 5'-untranslated region, a 1683-bp ORF, and a 36-bp 3'untranslated region. The ORF encodes a protein consisting of 560 amino acid residues and there existed seven putative 15 43. depicts transmembrane domains. Figure the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 56 kDa that was smaller than the molecular weight of 20 64,047 predicted from the ORF.

The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Mycobacterium tuberculosis hypothetical protein Rv0235c (Accession No. CAB07001).

Table 30 shows the comparison between amino acid sequences

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of the human protein of the present invention (HP) and Mycobacterium tuberculosis hypothetical protein Rv0235c (MT). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 41.7% in the entire region other than the N-terminal region. In addition, the region from alanine at position 293 to proline at position 502 matched with human putative novel protein c360B4.1 (Accession No. CAB56180).

Table 30

MT --AAVVAGAASFVPLW--ATMLIWLTLWVLYLSIVNVGQAWYSFGWESLLLETGFLMIFL

	HP	${\tt CPLWTLSRLPQHTPTSRIVLWGFRWLIFRIMLGAGLIKIRGDRCWRDLTCMDFHYETQPM}$
		.* .**. ***. ***. ***. ***. ***. ****. ****
	MT	GNERTAPPILTLLLA-RWLLFRVEFGAGLIKMRGDSCWRSLTCLYYHHETQPM
5		
	НР	PNPVAYYLHHSPWWFHRFETLSNHFIELLVPFFLFLGRRACIIHGVLQILFQAVLIVSGN
		*.*** * .**.*** * * ***
	MT	${\tt PGPLSWFFHHLPKPLHRIEVAGNHFAQLVVPFGLFTPQPAASIAAAIIVVTQLWLVASGN}$
10	НР	${\tt LSFLNWLTMVPSLACFDDATLGFLFPSGPGSLKDRVLQMQRDIRGARPEPRFGSVVRRAA}$
		.*.**** ******
	МТ	FSWLNWLTILLACSAIDTSS-AAALLPMPAQPALSAPPQWFAGLVV
	НР	${\tt NVSLGVLLAWLSVPVVLNLLSSRQVMNTHFNSLHIVNTYGAFGSITKERAEVILQGTASS}$
15		*** ** . *****.* ** ***.******** ***
	MT	VFTAAVLLLSYWPARNLLSSHQRMNMSFNPFHLVNTYGAFGSICRTRREVVIEGTDES
	HР	NASAPDAMWEDYEFKCKPGDPSRRPCLISPYHYRLDWLMWFAAFQTYEHNDWIIHLAGKL
20	MT	-PITEQTVWKAYEFKGKPGDPRRLPRQWAPYHLRLDWLMWFAAISPGYALPWMTPFLNRL
	HP	LASDAEALSLLAHNPFAGRPPPRWVRGEHYRYKFSRPGGRHAAEGKWWVRKRIGAYFPPL
		* . * * * * * * * * *
	MT	LRNDPATLKLLRHNPFP-QSPPRYVRAQLYQYRFTTVAELRRDRA-WWHRTLIGRYVPPM

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HP SLEELRPYFRDRGWPLPGPL

** ..

MT SLRKVASPPAD

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AA019047) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP03877> (SEQ ID NOS: 124, 134 and 144)

Determination of the whole base sequence of the cDNA insert of clone HP03877 obtained from cDNA library of 15 human kidney revealed the structure consisting of a 106-bp 5'-untranslated region, a 1221-bp ORF, and a 678-bp 3'untranslated region. The ORF encodes a protein consisting of 406 amino acid residues and there existed four putative 44 depicts the 20 transmembrane domains. Figure hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 49 kDa that was somewhat larger than the molecular weight 25 of 46,208 predicted from the ORF.

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The search of the protein database using the amino acid sequence of the present protein revealed that the protein was similar to Caenorhabditis elegans hypothetical protein Y37D8A (Accession No. CAA21543). Table 31 shows the comparison between amino acid sequences of the human protein of the present invention (HP) and Caenorhabditis elegans hypothetical protein Y37D8A (CE). Therein, the marks of -, *, and . represent a gap, an amino acid residue identical with that of the protein of the present invention, and an amino acid residue similar to that of the protein of the present invention, respectively. The both proteins shared a homology of 50.2% in the intermediate region of 329 amino acid residues.

15 Table 31

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HP MAENG

CE MAKKQKKSTEKSERTVEFKEPPKPANSEERLVSTRQFLAKIGQKKLIKKKVKNFRFSKKT

HP KNCDQRRVAMNKEHHNGNFTDPSSVNEKKRREREERQNIVLWRQPLITLQYFSLEILVIL
.* **..**. **..* .**..*

CE FIDFFSENQKKNCRLKPAGRGMKPSPSQNTLNRMERETIVFWRRPHIVIPYALMEIAHLA

HP KEWTSKLWHRQSIVVSFLLLLAVLIATYYVEGVHQQYVQRIEKQFLLYAYWIGLGILSSV

		* * *. *. ** ***. * *. *
	CE	VELFFKILAHKTVLLLTAISIGLAVYGYHAPGAHQEHVQTIEKHILWWSWWVLLGVLSSI
	НР	GLGTGLHTFLLYLGPHIASVTLAAYECNSVNFPEPPYPDQIICPDEEGTEGTISLWSIIS
5		***. ******. ******. **. **. **. * **
	CE	GLGSGLHTFLIYLGPHIAAVTMAAYECQSLDFPQPPYPESIQCPSTKSSI-AVTFWQIVA
	HР	KVRIEACMWGIGTAIGELPPYFMARAARLSGAEPDDEEYQEFEEMLEHAESAQDFA-
		. * ** ***. ****************
LO	CE	KVRVESLLWGAGTALGELPPYFMARAARISGQEPDDEEYREFLELMNADKESDADQKLSI
	НР	-SRAKLAVQKLVQKVGFFGILACASIPNPLFDLAGITCGHFLVPFWTFFGATLIGKAIIK
		·*** *** *** **********************
	CE	VERAKSWVEHNIHRLGFPGILLFASIPNPLFDLAGITCGHFLVPFWSFFGATLIGKALVK
L5		
	HP	MHIQKIFVIITFSKHIVEQMVAFIGAVPGIGPSLQKPFQEYLEAQRQKLHHKSEMGTPQG
		. *. *. **. * . *
	CE	MHVQMGFVILAFSDHHAENFVKILEKIPAVGPYIRQPISDLLEKQRKALHKTPGEHSEQD
20	HP	ENWLSWMFEKLVVVMVCYFILSIINSMAQSYAKRIQQRLNSEEKTK
	CE	LIDEENQSFEEEEEEAVTPPSSCPLLLSDGFEGVVVKK

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of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T18977) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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<HP10765> (SEQ ID NOS: 125, 135 and 145)

Determination of the whole base sequence of the cDNA insert of clone HP10765 obtained from cDNA library of human umbilical cord blood revealed the structure consisting of a 30-bp 5'-untranslated region, a 1362-bp ORF, and a 166-bp 3'-untranslated region. The ORF encodes a protein consisting of 453 amino acid residues and there existed a putative secretory signal at the N-terminus and one putative transmembrane domain in the inner portion. Figure 45 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 48 kDa that was almost identical with the molecular weight of 47,724 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI792834) among ESTs. However, since they are partial sequences, it can not be judged whether or not they

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encode the same protein as the protein of the present invention.

<HP10766> (SEQ ID NOS: 126, 136 and 146)

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Determination of the whole base sequence of the cDNA insert of clone HP10766 obtained from cDNA library of human kidney revealed the structure consisting of a 150-bp 5'-untranslated region, a 180-bp ORF, and a 675-bp 3'untranslated region. The ORF encodes a protein consisting of 59 amino acid residues and there existed two putative transmembrane domains. Figure 46 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 10 kDa or less that was almost identical with the molecular weight of 6,098 predicted from the ORF.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. T85491) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10770> (SEQ ID NOS: 127, 137 and 147)

Determination of the whole base sequence of the cDNA insert of clone HP10770 obtained from cDNA library of

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human kidney revealed the structure consisting of a 150-bp 5'-untranslated region, a 633-bp ORF, and a 186-bp 3'-untranslated region. The ORF encodes a protein consisting of 210 amino acid residues and there existed two putative transmembrane domains. Figure 47 depicts the hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of 27 kDa that was larger than the molecular weight of 22,156 predicted from the ORF.

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The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI792771) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10772> (SEQ ID NOS: 128, 138 and 148)

Determination of the whole base sequence of the

CDNA insert of clone HP10772 obtained from cDNA library of
human kidney revealed the structure consisting of a 19-bp
5'-untranslated region, a 498-bp ORF, and a 724-bp 3'untranslated region. The ORF encodes a protein consisting of
165 amino acid residues and there existed four putative
transmembrane domains. Figure 48 depicts the

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hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In vitro translation resulted in formation of a translation product of high molecular weight.

The search of the GenBank using the base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. F11871) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10773> (SEQ ID NOS: 129, 139 and 149)

Determination of the whole base sequence of the cDNA insert of clone HP10773 obtained from cDNA library of human kidney revealed the structure consisting of a 186-bp 5'-untranslated region, a 489-bp ORF, and a 499-bp 3'untranslated region. The ORF encodes a protein consisting of 162 amino acid residues and there existed four putative transmembrane domains. Figure 49 depicts hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. In translation resulted in formation of a translation product of high molecular weight.

The search of the GenBank using the base sequences

of the present cDNA has revealed the registration of

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sequences that shared a homology of 90% or more (for example, Accession No. N33828) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

<HP10776> (SEQ ID NOS: 130, 140 and 150) .

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Determination of the whole base sequence of the cDNA insert of clone HP10776 obtained from cDNA library of human kidney revealed the structure consisting of a 207-bp 5'-untranslated region, a 666-bp ORF, and a 139-bp 3'untranslated region. The ORF encodes a protein consisting of 221 amino acid residues and there existed three putative domains. Figure 50 transmembrane depicts hydrophobicity/hydrophilicity profile, obtained by the Kyte-Doolittle method, of the present protein. translation resulted in formation of a translation product of 30 kDa that was larger than the molecular weight of 24,883 predicted from the ORF.

base sequences of the present cDNA has revealed the registration of sequences that shared a homology of 90% or more (for example, Accession No. AI929639) among ESTs. However, since they are partial sequences, it can not be judged whether or not they encode the same protein as the protein of the present invention.

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INDUSTRIAL APPLICABILITY

The present invention provides human proteins having hydrophobic domains, DNAs encoding these proteins, expression vectors for these DNAs and eukaryotic cells 5 expressing these DNAs. Since all of the proteins of the present invention are secreted or exist in the cell membrane, are considered to be proteins controlling they proliferation and/or the differentiation of the cells. Accordingly, the proteins of the present invention can be 10 employed as pharmaceuticals such as carcinostatic agents which act to control the proliferation and/or differentiation of the cells, or as antigens for preparing antibodies against these proteins. The DNAs of the present invention can be utilized as probes for the genetic 15 diagnosis and gene sources for the gene therapy. Furthermore, the DNAs can be utilized for expressing these proteins in large quantities. Cells into which these genes introduced to express these proteins can be utilized for 20 detection of the corresponding receptors or ligands, screening of novel small molecule pharmaceuticals and the like. The antibody of the present invention can be utilized for the detection, quantification, purification and the like of the protein of the present invention.

The present invention also provides genes

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corresponding to the polynucleotide sequences disclosed herein. "Corresponding genes" are the regions of the genome that are transcribed to produce the mRNAs from which cDNA polynucleotide sequences are derived and may include contiguous regions of the genome necessary for the regulated expression of such genes. Corresponding genes may therefore include but are not limited to coding sequences, 5' and 3' untranslated regions, alternatively spliced exons, introns, promoters, enhancers, and silencer or suppressor elements. The corresponding genes can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include the preparation of probes or the disclosed sequence primers from information identification and/or amplification of genes in appropriate genomic libraries or other sources of genomic materials. An "isolated gene" is a gene that has been separated from the adjacent coding sequences, if any, present in the genome of the organism from which the gene was isolated.

organisms that have enhanced, reduced, or modified expression of the gene(s) corresponding to the polynucleotide sequences disclosed herein are provided. The desired change in gene expression can be achieved through the use of antisense polynucleotides or ribozymes that bind and/or cleave the mRNA transcribed from the gene (Albert and Morris, 1994, Trends Pharmacol. Sci. 15(7): 250-254;

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Lavarosky et al., 1997, Biochem. Mol. Med. 62(1): 11-22; and Hampel, 1998, Prog. Nucleic Acid Res. Mol. Biol. 58: 1-39; all of which are incorporated by reference herein). Transgenic animals that have multiple copies of the gene(s) corresponding to the polynucleotide sequences disclosed herein, preferably produced by transformation of cells with genetic constructs that are stably maintained within the transformed cells and their progeny, are provided. Transgenic animals that have modified genetic control regions that increase or reduce gene expression levels, or that change temporal or spatial patterns of gene expression, are also provided (see European Patent No. 0 649 464 Bl, incorporated by reference herein), In addition, organisms are provided in which the gene(s) corresponding to the polynucleotide sequences disclosed herein have been partially or completely inactivated, through insertion of extraneous sequences into the corresponding gene(s) through deletion of all or part of the corresponding gene(s). Partial or complete gene inactivation can be accomplished through insertion, preferably followed by imprecise excision, of transposable elements (Plasterk, 1992, Bioessays 14(9): 629-633; Zwaal et al., 1993, Proc. Natl. Acad. Sci. USA 90(16): 7431-7435; Clark et al., 1994, Proc. Natl. Acad. Sci. USA 91(2): 719-722; all of which are incorporated by reference herein), or through homologous recombination,

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preferably detected by positive/negative genetic selection strategies (Mansour et al., 1988, Nature 336: 348-352; U.S. Patent Nos. 5,464,764; 5,487,992; 5,627,059; 5,631,153; 5,614, 396; 5,616,491; and 5,679,523; all of which are incorporated by reference herein). These organisms with altered gene expression are preferably eukaryotes and more preferably are mammals. Such organisms are useful for the development of non-human models for the study of disorders involving the corresponding gene(s), and for the development of assay systems for the identification of molecules that interact with the protein product(s) of the corresponding gene(s). Where the protein of the present invention is membrane-bound (e.g., is a receptor), the present invention also provides for soluble forms of such protein. In such forms part or all of the intracellular and transmembrane domains of the protein are deleted such that the protein is fully secreted from the cell in which it is expressed. The intracellular and transmembrane domains of proteins of the identified in accordance with known invention can be techniques for determination of such domains from sequence information.

Proteins and protein fragments of the present invention include proteins with amino acid sequence lengths that are at least 25% (more preferably at least 50%, and most preferably at least 75%) of the length of a disclosed

protein and have at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% or 95% identity) with that disclosed protein, where sequence identity is determined by comparing the amino acid sequences of the proteins when aligned so as to maximize overlap and identity while minimizing sequence gaps. Also included in the present invention are proteins and protein fragments that contain a segment preferably comprising 8 or more (more preferably 20 or more, most preferably 30 or more) contiguous amino acids that shares at least 75% sequence identity (more preferably, at least 85% identity; most preferably at least 95% identity) with any such segment of any of the disclosed proteins.

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Species homologs of the disclosed polynucleotides and proteins are also provided by the present invention. As used herein, a "species homologue" is a protein or polynucleotide with a different species of origin from that of a given protein or polynucleotide, but with significant sequence similarity to the given protein or polynucleotide, as determined by those of skill in the art. Species homologs may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source from the desired species.

The invention also encompasses allelic variants of the disclosed polynucleotides or proteins; that is,

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naturally-occurring alternative forms of the isolated polynucleotide which also encode proteins which are identical, homologous, or related to that encoded by the polynucleotides.

The invention also includes polynucleotides with sequences complementary to those of the polynucleotides disclosed herein.

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The present invention also includes polynucleotides capable of hybridizing under reduced stringency conditions, more preferably stringent conditions, and most preferably highly stringent conditions, to polynucleotides described herein. Examples of stringency conditions are shown in the table below: highly stringent conditions are those that are at least as stringent as, for example, conditions A-F; stringent conditions are at least as stringent as, for example, conditions are at least as stringent as, for example, conditions are at least as stringent as, for example, conditions G-L; and reduced stringency conditions are at least as stringent as, for example, conditions M-R.

Table 32

Stringency	Poly-	Hybrid	Hybridization Temperature	Wash
Condition	nucleotide	Length	and Buffer'	Temperature
	Hybrid	(bp) '		and Buffer'
A	DNA: DNA	≥50	65°C; 1×SSC -or-	65°C;
			42°C; 1×SSC,50%	0.3×SSC
			formamide	
В	DNA: DNA	<50	T _B *; 1×SSC	T _B *; 1×SSC
С	DNA: RNA	≥50	67°C; 1×SSC -or-	67°C;
			45°C; 1×SSC,50%	0.3×SSC
			formamide	
D	DNA: RNA	<50	T _b *; 1×SSC	To*; 1×SSC
E	RNA: RNA	≥50	70°C; 1×SSC -or-	70°C;
İ			50°C; 1×SSC,50%	0.3×SSC
			formamide	
F	RNA: RNA	<50	T _F *; 1×SSC	T _F *; 1×SSC
G	DNA: DNA	≥50	65°C; 4×SSC -or-	65°C; 1×SSC
			42°C; 4×SSC,50%	
			formamide	
Н	DNA: DNA	<50	T _H *; 4×SSC	T _H *; 4×SSC
I	DNA: RNA	≥50	67°C; 4×SSC -or-	67°C; 1×SSC
			45°C; 4×SSC,50%	
			formamide	
J	DNA: RNA	<50	T _J *; 4×SSC	T _J *; 4×SSC
K	RNA: RNA	≥50	70°C; 4×SSC -or-	67°C; 1×SSC
		1	50°C; 4×SSC,50%	
			formamide	
L	RNA: RNA	<50	T _L *; 2×SSC	T _L *; 2×SSC
M	DNA: DNA	≥50	50°C; 4×SSC -or-	50°C; 2×SSC
			40°C; 6×SSC,50%	
			formamide	
N	DNA: DNA	<50	T _N *; 6×SSC	T _N *; 6×SSC
0	DNA: RNA	≥50	55°C; 4×SSC -or-	55°C; 2×SSC
			42°C; 6×SSC,50%	
	<u> </u>		formamide	
P	DNA: RNA	<50	T,*; 6×SSC	Tp*; 6×SSC
Q	RNA: RNA	≥50	60°C; 4×SSC -or-	60°C; 2×SSC
			45°C; 6×SSC,50%	1
	ļ,	<u> </u>	formamide	<u> </u>
R	RNA: RNA	<50	T _R *; 4×SSC	T _R *; 4×SSC

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- ‡ : The hybrid length is that anticipated for the hybridized region(s) of the hybridizing polynucleotides.

 When hybridizing a polynucleotide to a target polynucleotide of unknown sequence, the hybrid length is assumed to be that of the hybridizing polynucleotide. When polynucleotides of known sequence are hybridized, the hybrid length can be determined by aligning the sequences of the polynucleotides and identifying the region or regions of optimal sequence complementarity.
- † : SSPE (1×SSPE is 0.15M NaCl, 10mM NaH₂PO₄, and 1.25mM EDTA, pH7.4) can be substituted for SSC (1×SSC is 0.15M NaCl and 15mM sodium citrate) in the hybridization and wash buffers; washes are performed for 15 minutes after hybridization is complete.
- *T_B T_R: The hybridization temperature for hybrids
 anticipated to be less than 50 base pairs in length should
 be 5-10°C less than the melting temperature (T_m) of the
 hybrid, where T_m is determined according to the following
 equations. For hybrids less than 18 base pairs in length,

 T_m(°C)=2(#of A + T bases) + 4(# of G + C bases). For hybrids
 between 18 and 49 base pairs in length, T_m(°C)=81.5 +
 16.6(log₁₀[Na⁺]) + 0.41 (%G+C) (600/N), where N is the
 number of bases in the hybrid, and [Na⁺] is the concentration
 of sodium ions in the hybridization buffer ([Na⁺] for
 1×SSC=0.165M).

Additional examples of stringency conditions for polynucleotide hybridization are provided in Sambrook, J., E.F. Fritsch, and T. Maniatis, 1989, Molecular Cloning: A Laboratory Manual, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, chapters 9 and 11, and Current Protocols in Molecular Biology, 1995, F.M. Ausubel et al., eds., John Wiley & Sons, Inc., sections 2.10 and 6.3-6.4, incorporated herein by reference.

Preferably, each such hybridizing polynucleotide has a length that is at least 25% (more preferably at least 50%, and most preferably at least 75%) of the length of the polynucleotide of the present invention to which it hybridizes, and has at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% or 95% identity) with the polynucleotide of the present invention to which it hybridizes, where sequence identity is determined by comparing the sequences of the hybridizing polynucleotides when aligned so as to maximize overlap and identity while minimizing sequence gaps.

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CLAIMS

1. A protein comprising any one of an amino acid sequence selected from the group consisting of SEQ ID NOS: 1 to 10, 31 to 40, 61 to 70, 91 to 100 and 121 to 130.

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- 2. An isolated DNA encoding the protein according to Claim 1.
- 3. An isolated cDNA comprising any one of a base sequence selected from the group consisting of SEQ ID NOS: 11 to 20, 41 to 50, 71 to 80, 101 to 110 and 131 to 140.
- 4. The cDNA according to Claim 3 consisting of any one of a base sequence selected from the group consisting of SEQ ID NOS: 21 to 30, 51 to 60, 81 to 90, 111 to 120 and 141 to 150.
- 5. An expression vector that is capable of expressing the DNA according to any one of Claim 2 to Claim 4 by in vitro translation or in eukaryotic cells.
 - 6. A transformed eukaryotic cell that is capable of expressing the DNA according to any one of Claim 2 to Claim 4 and of producing the protein according to Claim 1.
 - 7. An antibody directed to the protein according to Claim 1.

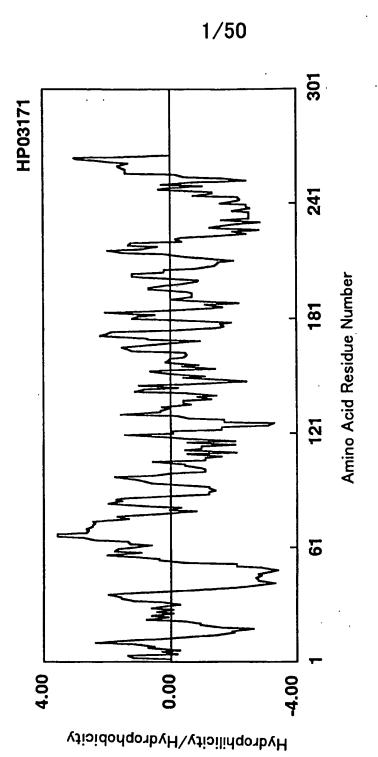


Fig.1



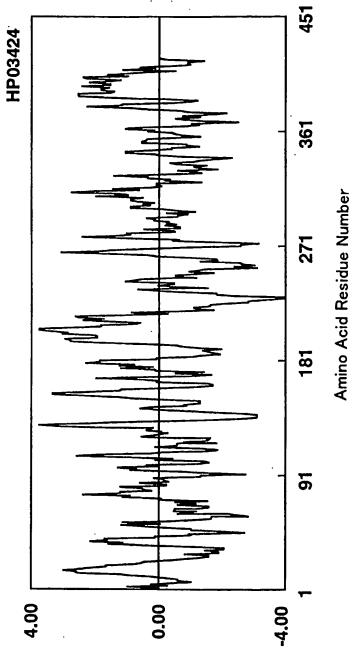


Fig.2

Hydrophilicity/Hydrophobicity

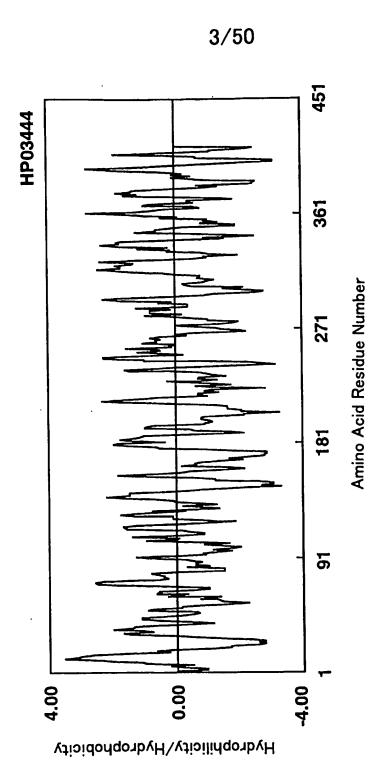
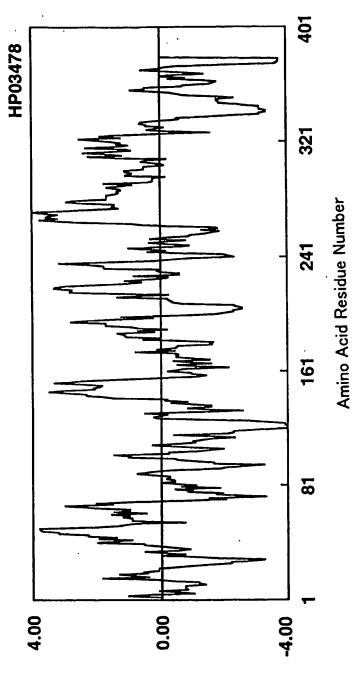


Fig.3





Hydrophilicity/Hydrophobicity

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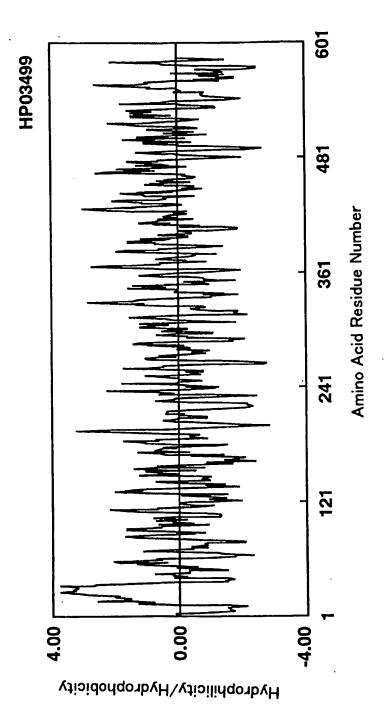
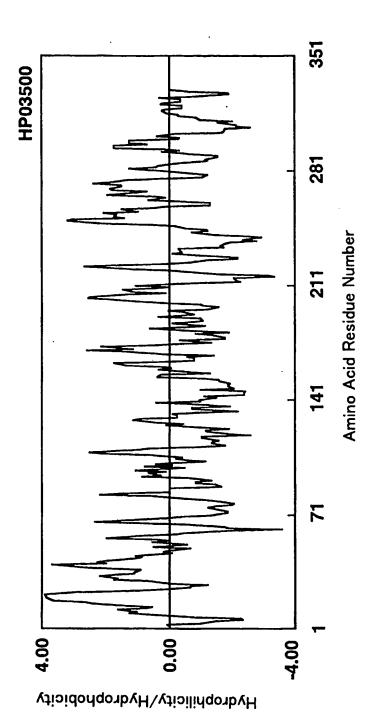


Fig.5





F18.6

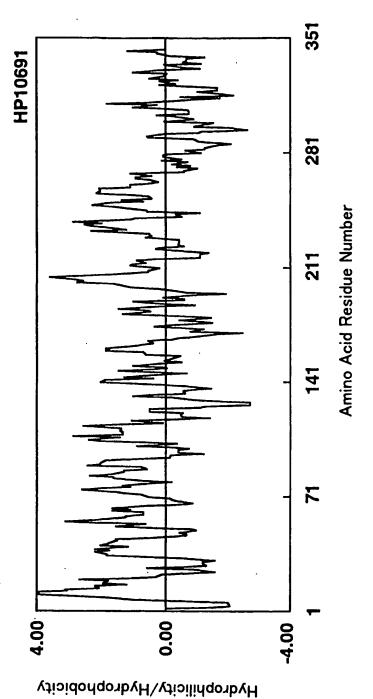


Fig. 7



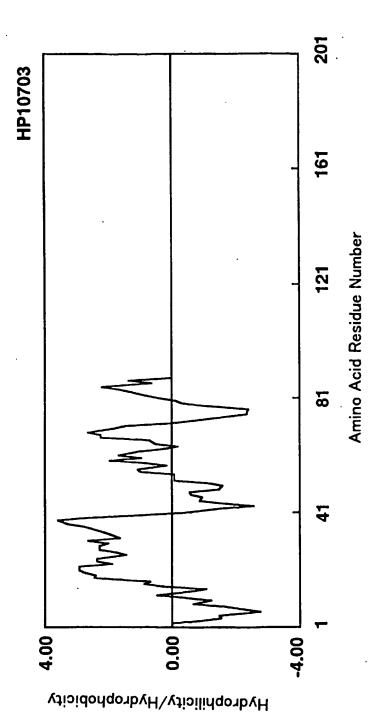


Fig.8



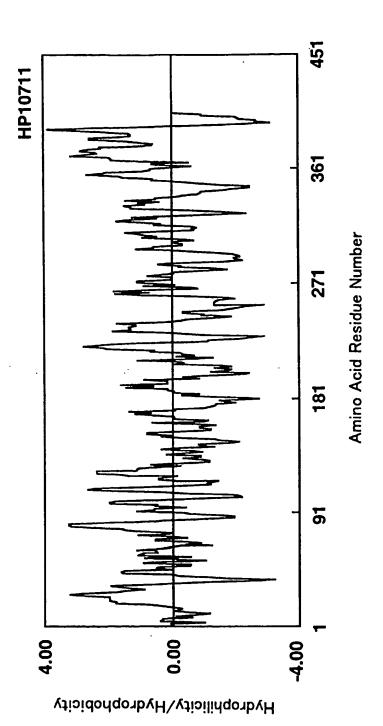
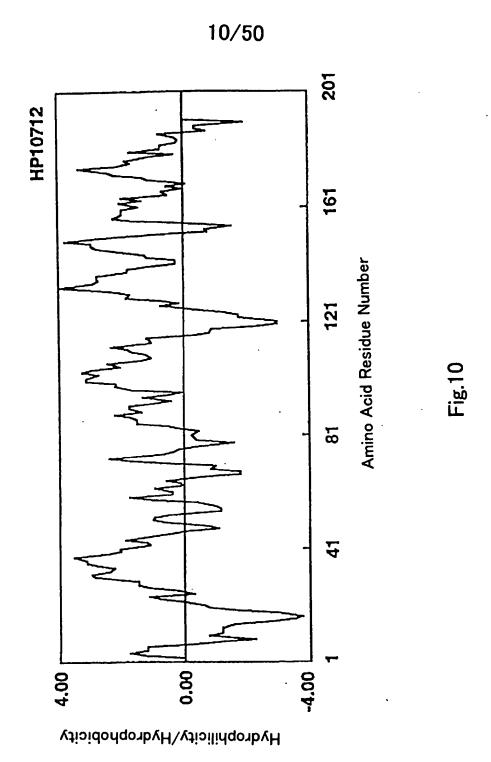


Fig.9





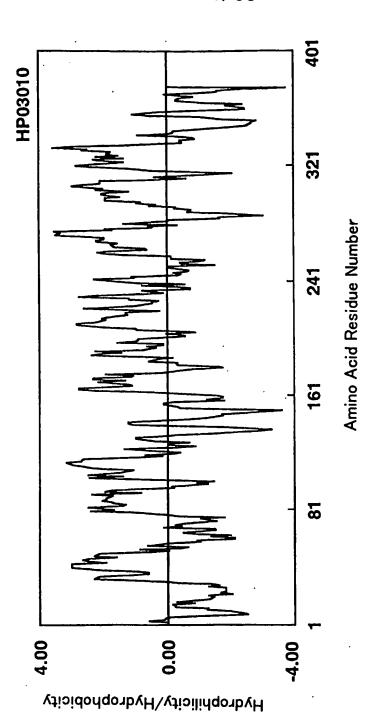


Fig. 1.1



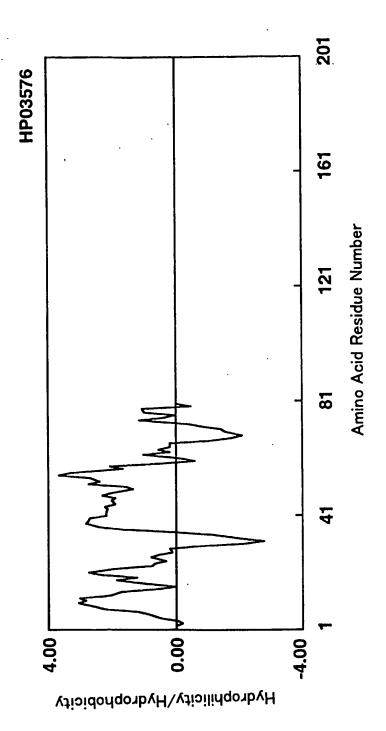


Fig. 12



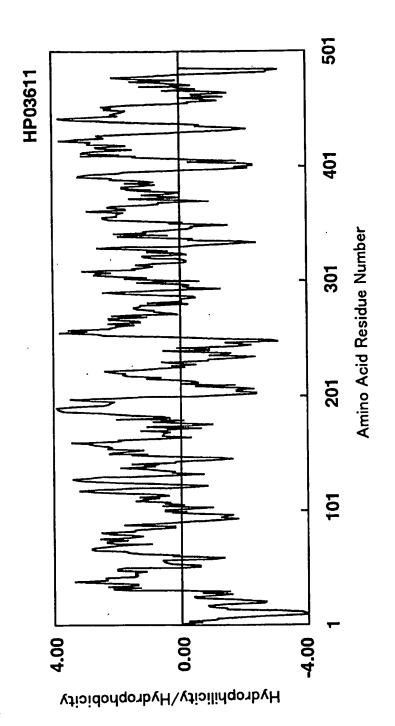


Fig.13



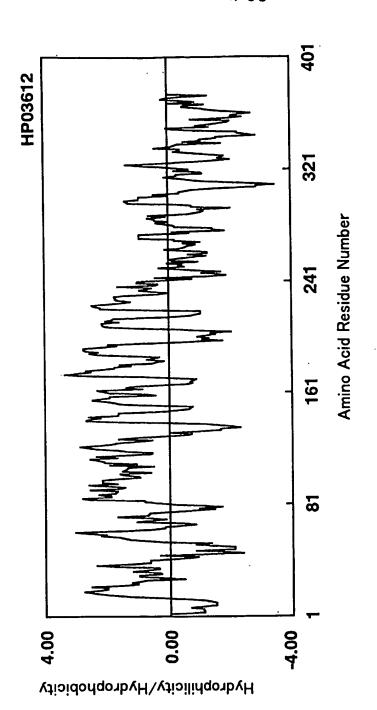


Fig. 14



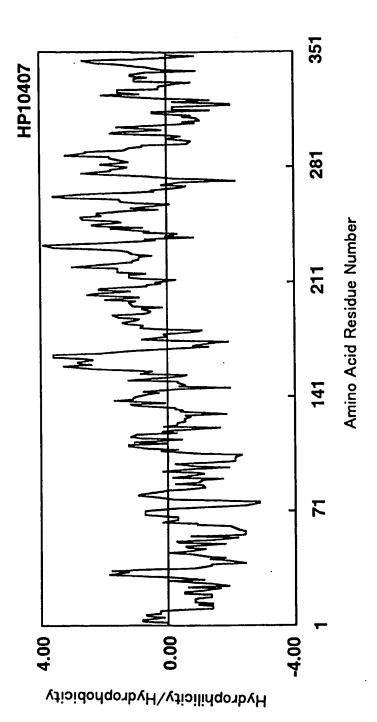


Fig.15



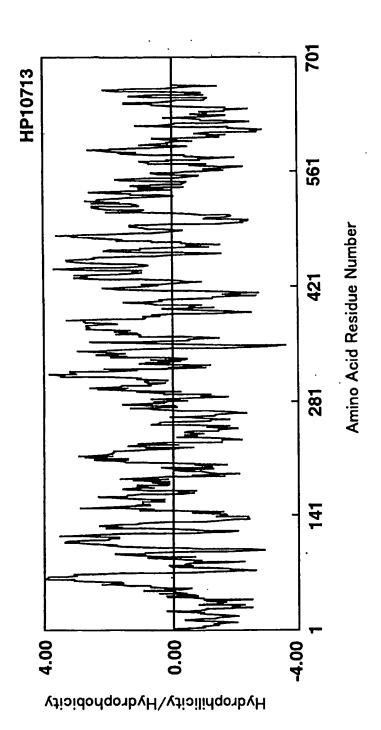
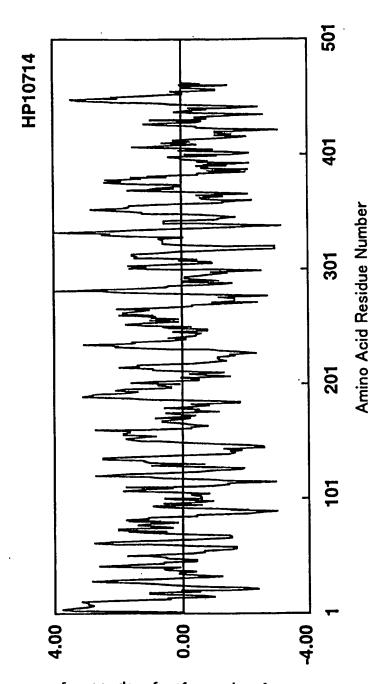


Fig. 16





Hydrophilicity/Hydrophobicity

Fig.17

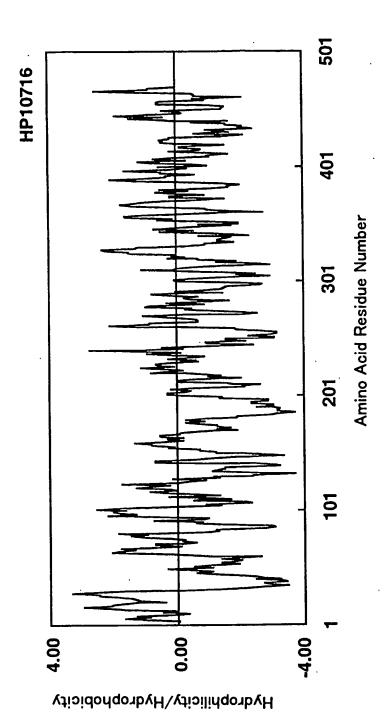


Fig. 18



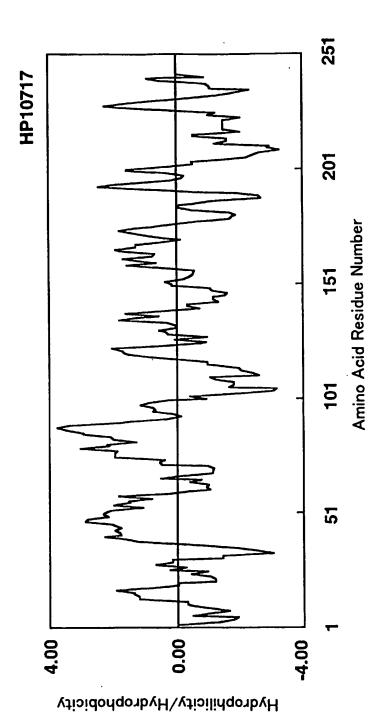
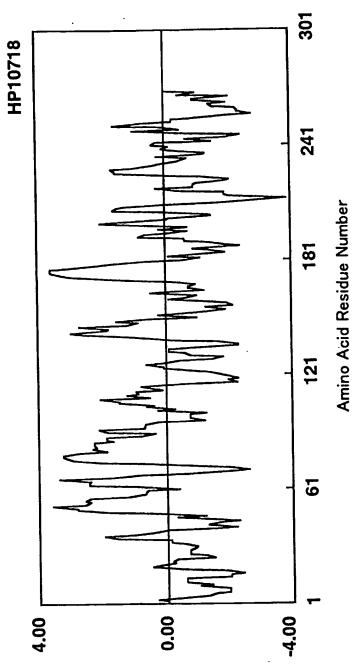


Fig. 19





Hydrophilicity/Hydrophobicity

1g.20



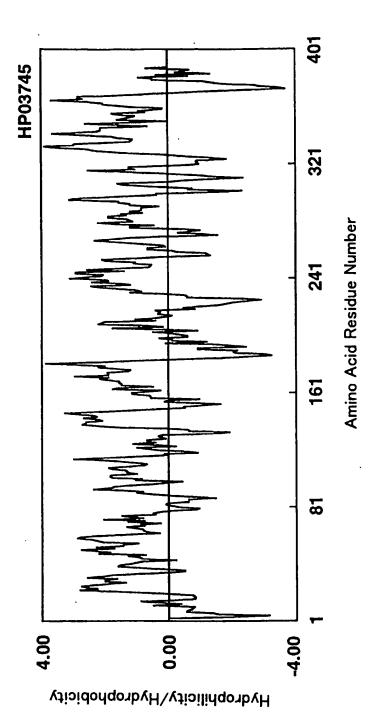
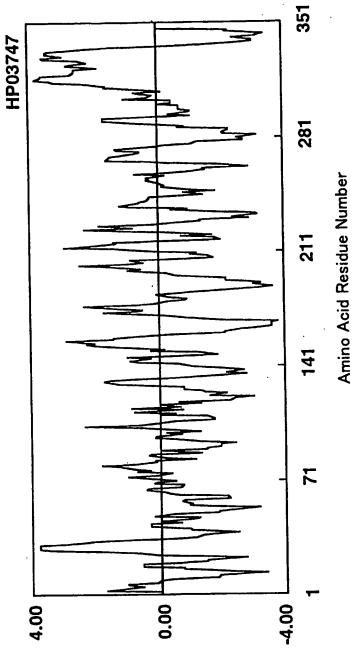


Fig.21



- Hydrophilicity/Hydrophobicity

-ig.22



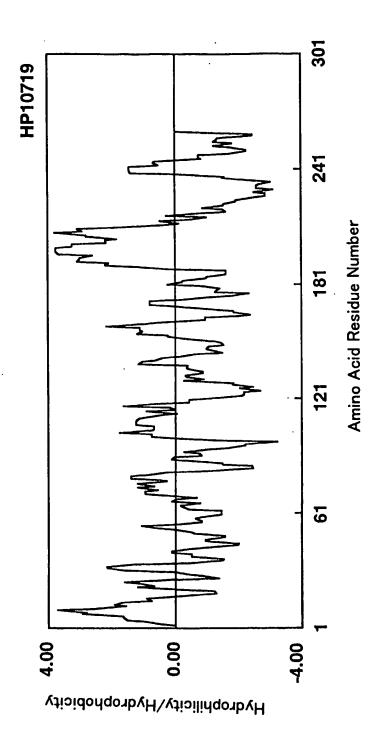


Fig. 23

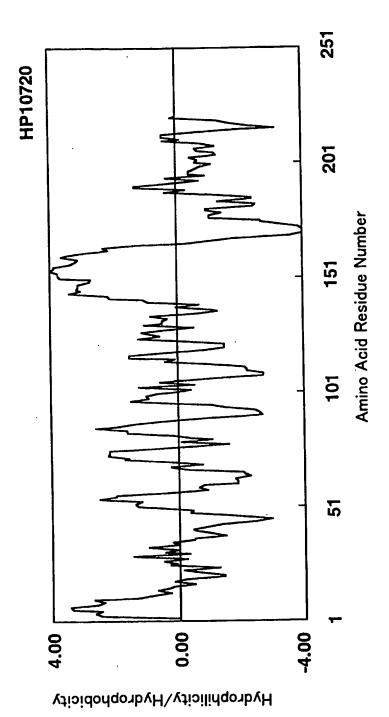


Fig.24



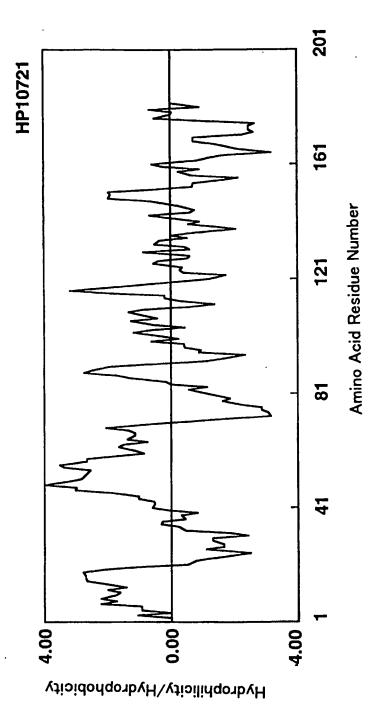


Fig.25



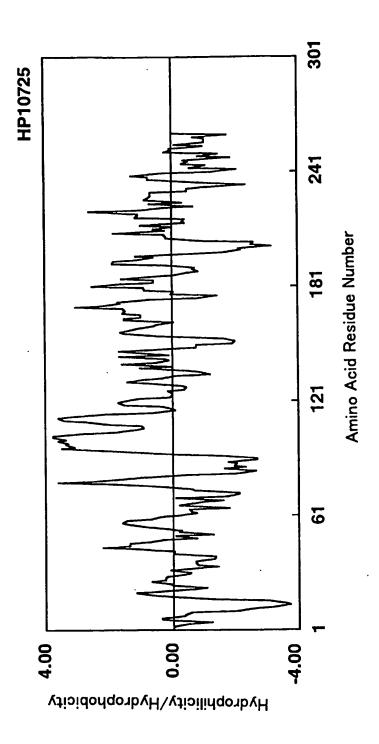


Fig.26



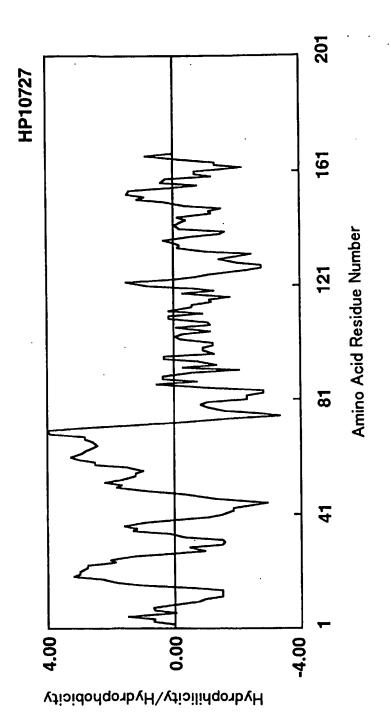
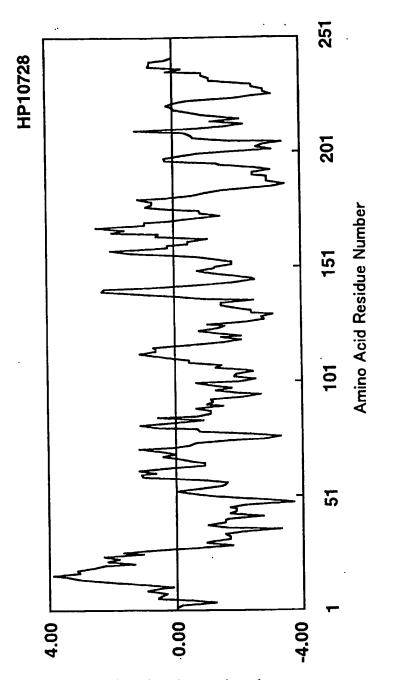


Fig.2





ΗλακορλίΙισίτη/Ηγακορλορίσίτη.

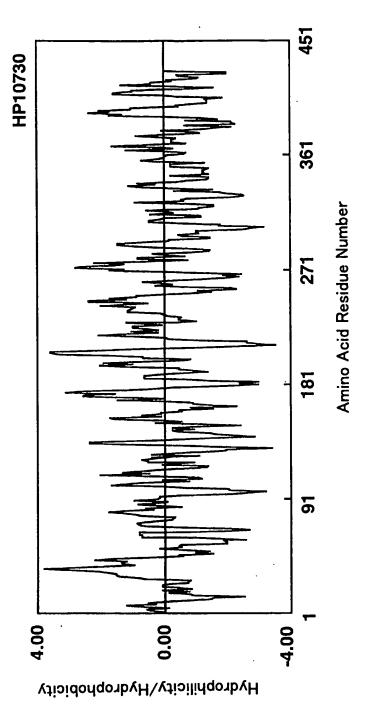


Fig.29

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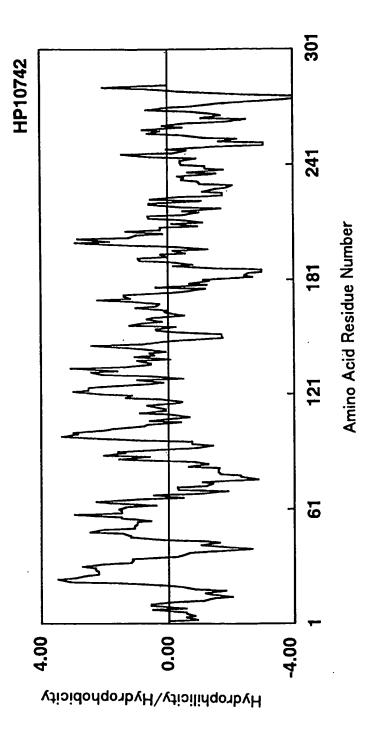
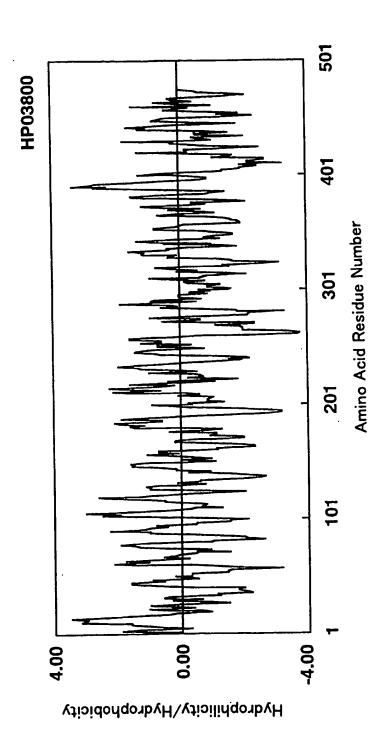


Fig. 30





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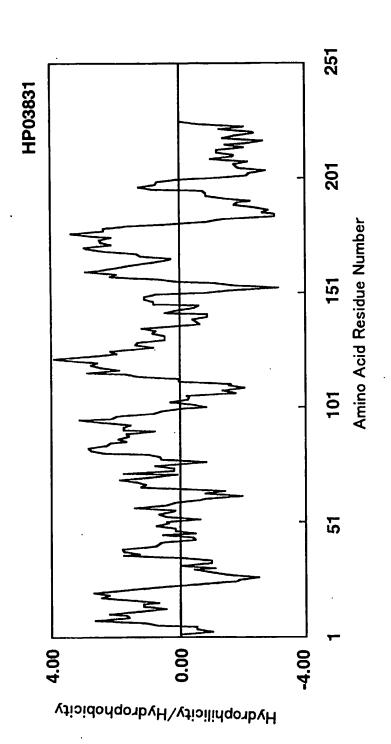


Fig. 32



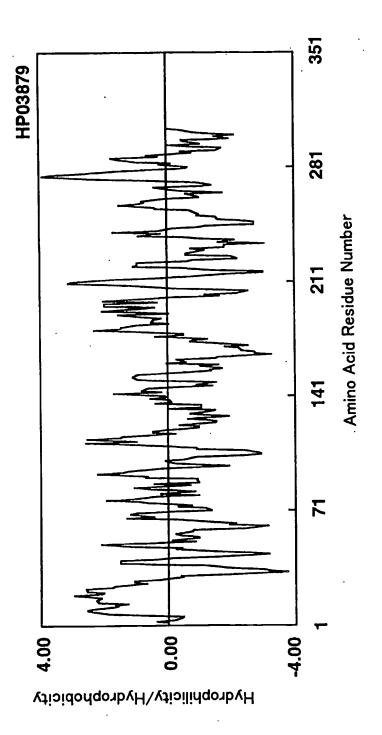


Fig.3

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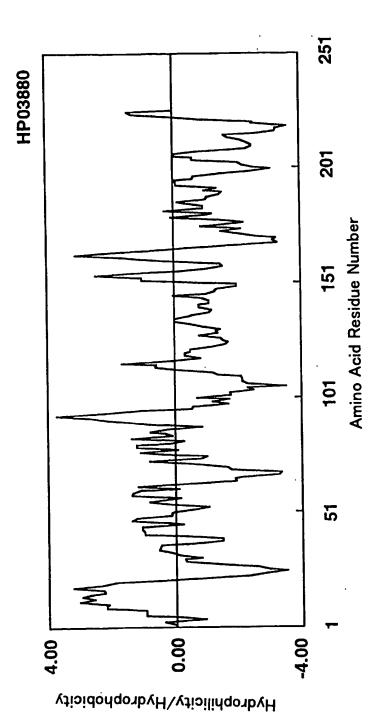


Fig.34

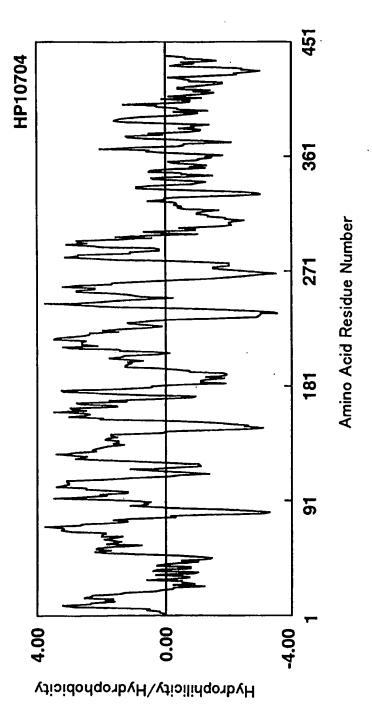


Fig.35

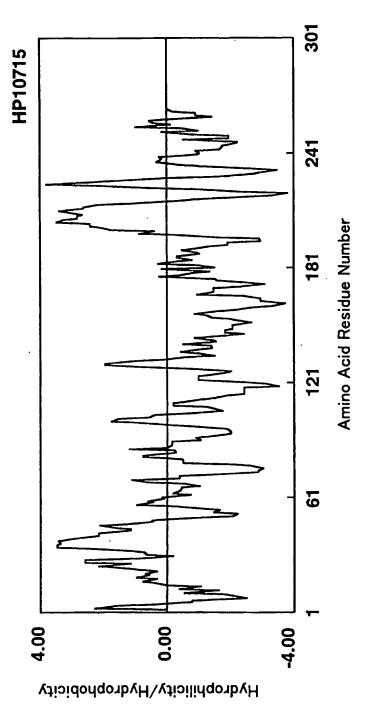


Fig.36

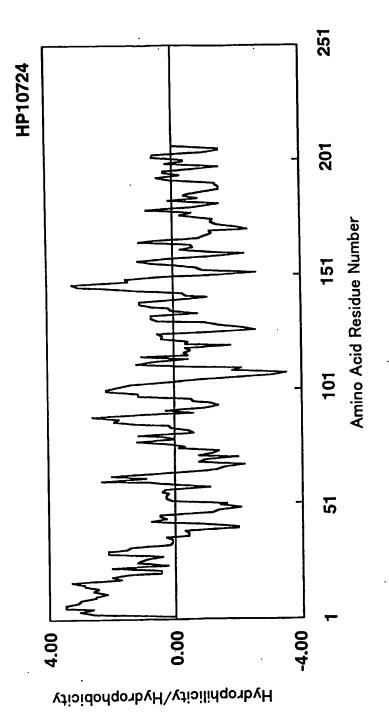


Fig. 37

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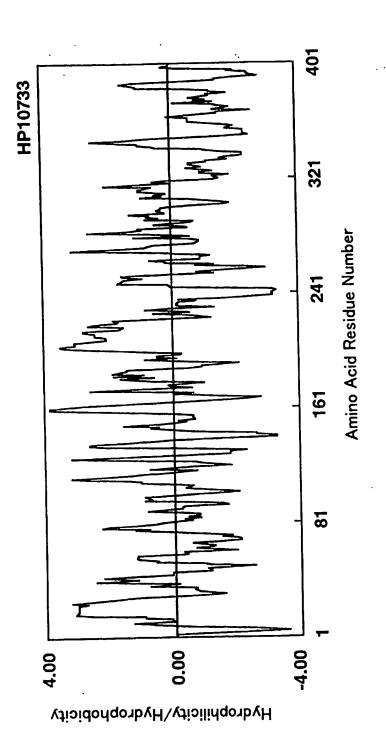


Fig.38



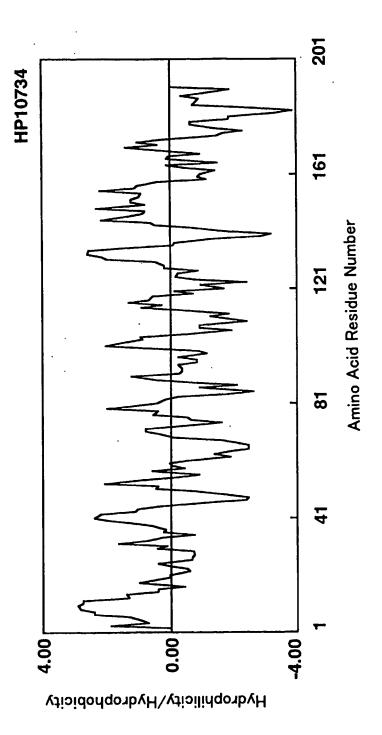


Fig. 39



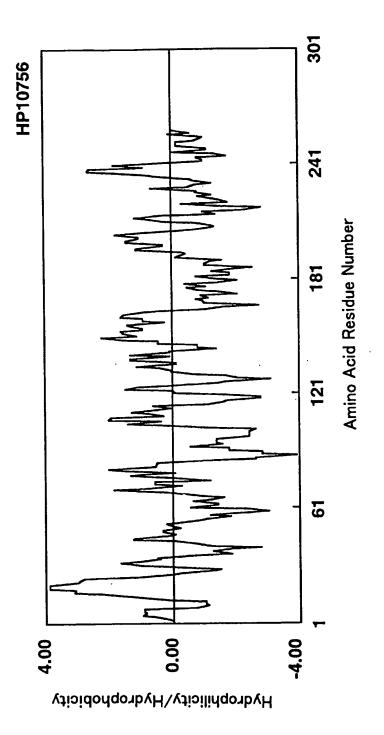


Fig.40

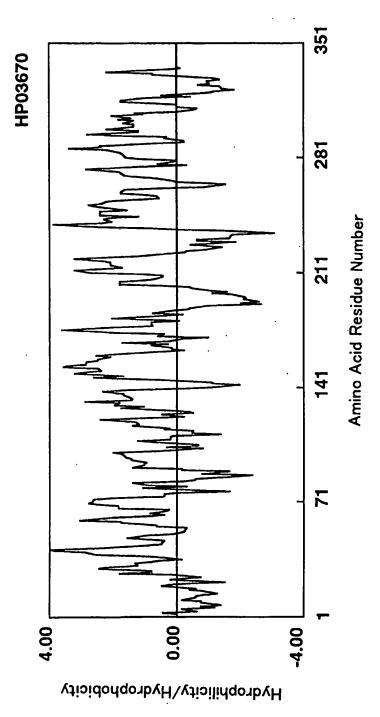


Fig.41

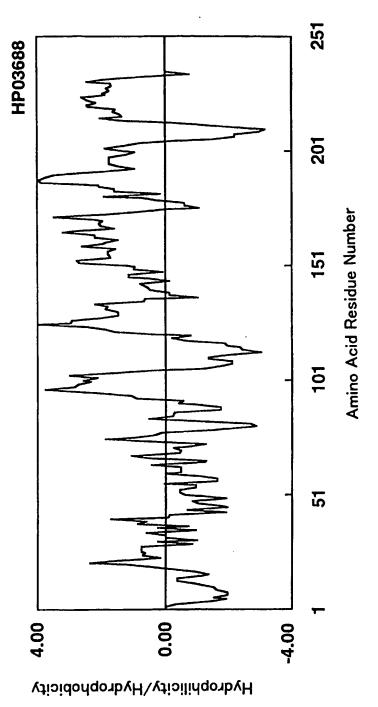


Fig. 42



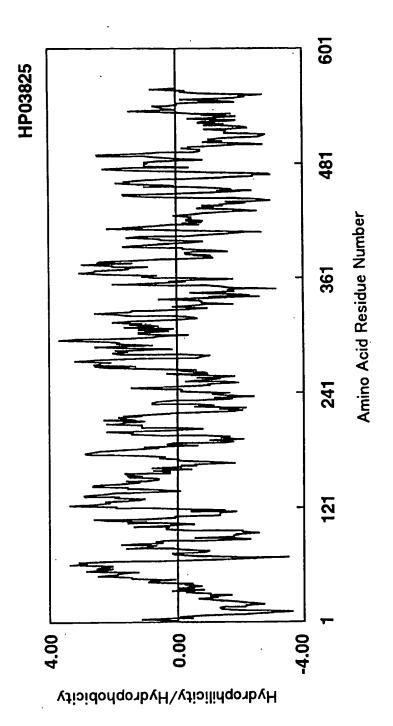


Fig.43

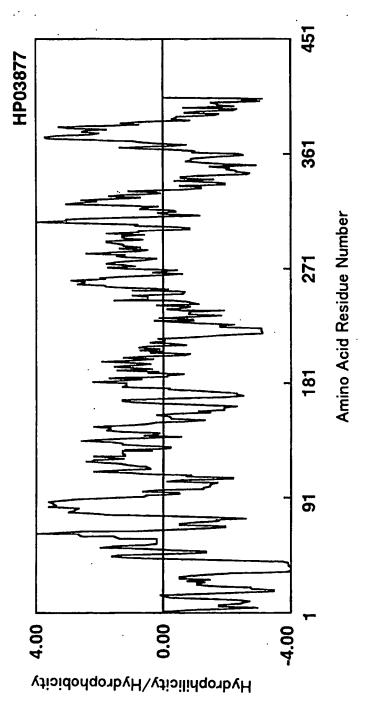


Fig.44



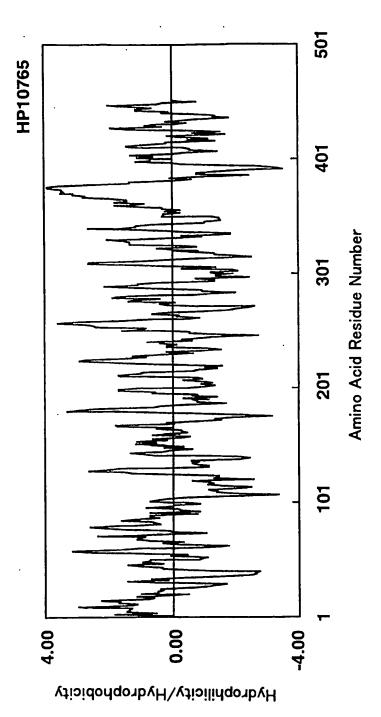


Fig. 45



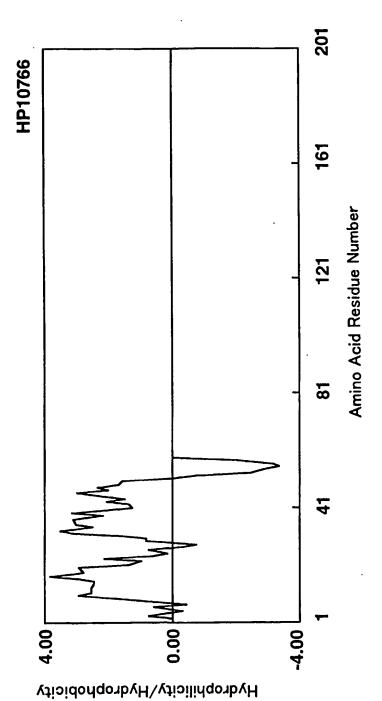


Fig.46



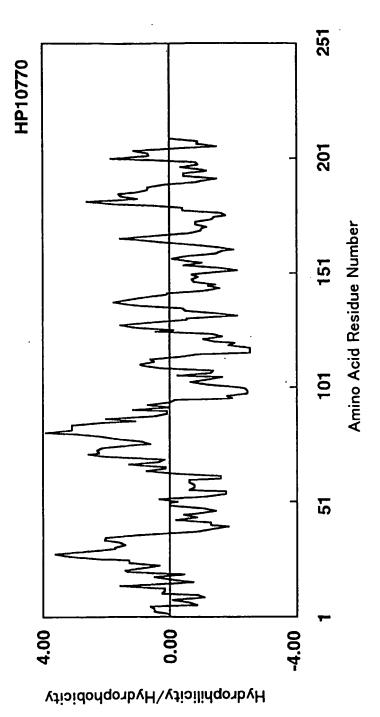


Fig.47



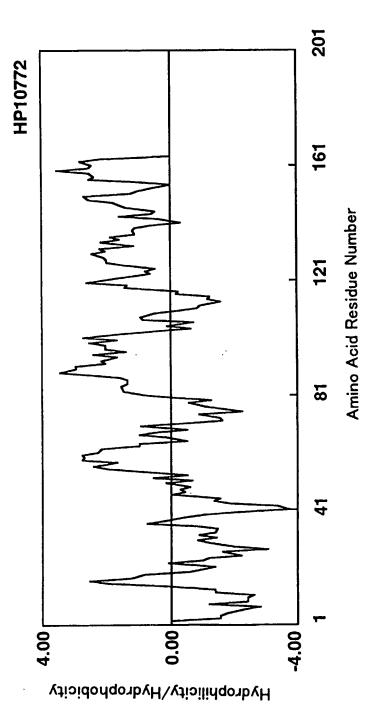


Fig.48



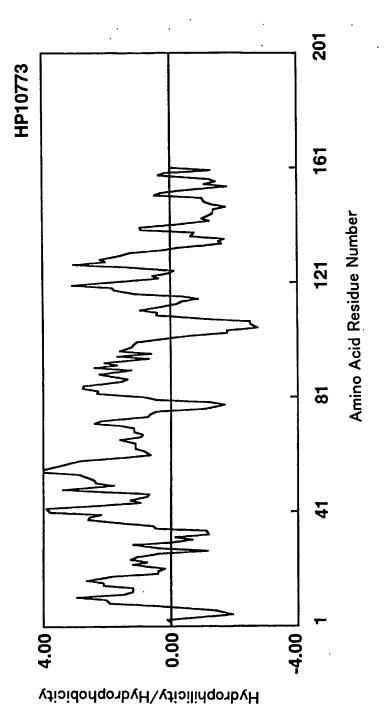


Fig. 49



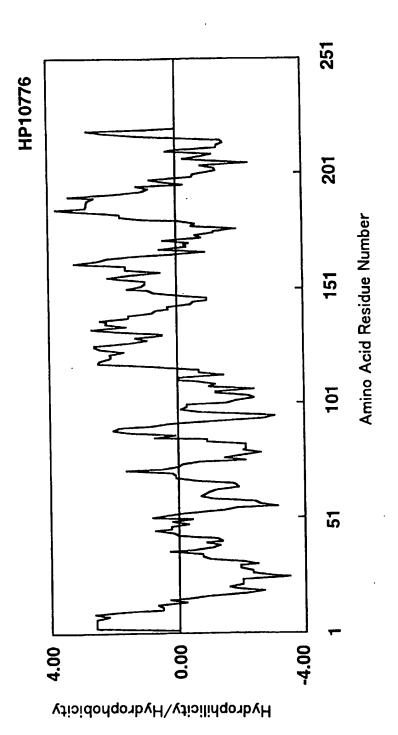


Fig.50

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Protegene Inc.

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Glu Ile Leu Leu Thr Pro Ala Arg Glu Glu Gln Pro Pro Gln His Arg

35 40 45

Ser Lys Arg Gly Ser Ser Val Gly Gly Val Cys Tyr Leu Ser Met Gly

50 55 60

Met Val Val Leu Leu Met Gly Leu Val Phe Ala Ser Val Tyr Ile Tyr

65 70 75 80

Arg Tyr Phe Phe Leu Ala Gln Leu Ala Arg Asp Asn Phe Phe Arg Cys

85 90 95

Gly Val Leu Tyr Glu Asp Ser Leu Ser Ser Gln Val Arg Thr Gln Met

100 105 110

Glu Leu Glu Glu Asp Val Lys Ile Tyr Leu Asp Glu Asn Tyr Glu Arg

115 120 125

Ile Asn Val Pro Val Pro Gln Phe Gly Gly Gly Asp Pro Ala Asp Ile

130 135 140

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Ile His Asp Phe Gln Arg Gly Leu Thr Ala Tyr His Asp Ile Ser Leu Asp Lys Cys Tyr Val Ile Glu Leu Asn Thr Thr Ile Val Leu Pro Pro Arg Asn Phe Trp Glu Leu Leu Met Asn Val Lys Arg Gly Thr Tyr Leu Pro Gln Thr Tyr Ile Ile Gln Glu Glu Met Val Val Thr Glu His Val Ser Asp Lys Glu Ala Leu Gly Ser Phe Ile Tyr His Leu Cys Asn Gly Lys Asp Thr Tyr Arg Leu Arg Arg Arg Ala Thr Arg Arg Arg Ile Asn Lys Arg Gly Ala Lys Asn Cys Asn Ala Ile Arg His Phe Glu Asn Thr Phe Val Val Glu Thr Leu Ile Cys Gly Val Val ⟨210⟩ 2 <211> 419 <212> PRT <213> Homo sapiens <400> 2 Met Ser Cys Ala Gly Arg Ala Gly Pro Ala Arg Leu Ala Ala Leu Ala Leu Leu Thr Cys Ser Leu Trp Pro Ala Arg Ala Asp Asn Ala Ser Gln

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Gly	Ala	Pro	Leu	Thr	Phe	Arg	Ile	Asp	Arg	Gly	Arg	Tyr	Gly	Leu	Asp
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Ser	Pro	Lys	Ala	Glu	Val	Arg	Gly	Gln	Val	Leu	Ala	Pro	Leu	Pro	Leu
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His	Gly	Val	Ala	Asp	His	Leu	Gly	Cys	Asp	Pro	Gln	Thr	Arg	Phe	Phe
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Cys	Thr	Phe	Lys	Glu	Lys	Ile	Ser	Arg	Ala	Ala	Phe	His	Asn	Ala	Val
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Thr	His	Pro	Gly	Thr	Gly	Asp	Ile	Ile	Ala	Val	Met	Ile	Thr	Glu	Leu
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Arg	Gly	Lys	Asp	Ile	Leu	Ser	Tyr	Leu	Glu	Lys	Asn	Ile	Ser	Val	Gln
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A	410	A	Aan	Arg	Aon	Gl n	Ara	Ara	וום 1	G1 v	Asn	Ala	Ala	l.vs	Lvs
ASII	VIA	ив	ASP	ΜŘ		GIII	мв	шБ	Dea		nop			_,_	
225					230					235					240
Ala	Ile	Ser	Lys	Leu	Thr	Thr	Arg	Thr	Val	Lys	Lys	Gly	Asp	Lys	Glu
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Thr	Asp	Pro	Asp	Phe	Asp	His	Cys	Ala	Val	Cys	Ile	Glu	Ser	Tyr	Lys
			260					265					270		
G1n	Aen	Asn		Val	Arg	Ile	Leu	Pro	Cvs	Lvs	His	Val	Phe	His	Lys
0111	11311		,,,	,,,,			280		.,.	-,-		285			
		275			_	_			•••	•	6 0		D	W- +	Cons
Ser	Cys	Val	Asp	Pro	Trp	Leu	Ser	Glu	His	Cys	Ihr	Cys	Pro	Met	cys
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Lys	Leu	Asn	Ile	Leu	Lys	Ala	Leu	Gly	Ile	Val	Pro	Asn	Leu	Pro	Cys
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Thr	Asp	Asn	Val	Ala	Phe	Asp	Met	Glu	Arg	Leu	Thr	Arg	Thr	G1n	Ala
				325					330					335	
Va 1	Asn	Aro	Aro	Ser	Ala	Leu	Glv	Asp	Leu	Ala	Gly	Asp	Asn	Ser	Leu
101	11011	6					,	345			•	-	350		
			340							•	_	•		C1	A
Gly	Leu	Glu	Pro	Leu	Arg	Thr	Ser	Gly	lle	Ser	Pro			Gin	ASP
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Gly	Glu	ı Let	Thi	Pro	Arg	Thr	Gly	Glu	Ile	Asn	Ile	Ala	Val	Thr	Lys
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Gli	ı Trp	Phe	e Ile	e Ile	e Ala	Ser	Phe	Gly	Leu	Leu	Ser	· Ala	Leu	Thr	Leu
385	5				390)				395	j				400
		r Mas	+ 114	o 114			: Thr	- A1s	. Set	Ler	ı Asr	n Ala	ı Asr	Glu	Val
Oy:	. 131	, nic	, 11,			,								415	
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Glı	ı Tr	p Ph	е												

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Cys	Gly	Gly	Ile	Leu	Thr	Gly	Glu	Ser	Gly	Phe	Ile	Gly	Ser	Glu	Gly
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Thr	Ala	Asp	Gly	Phe	Ile	Gly	His	Tyr	Ile	Phe	Arg	Pro	Lys	Lys	Leu
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Thr	Val	Ile	Thr	Thr	Ile	Thr	Arg	Asp	Gly	Ser	Leu	His	Ala	Thr	Val
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Ser	Ile	Ile	Asn	Ile	Tyr	Lys	Glu	Gly	Asn	Leu	Ala	Ile	Gln	G1n	Ala
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Pro	Val	Asr	Leu	ı Thr	Trp	Ala	Asp	Leu	Glu	Asp	Arg	Asp	Gly	Arg	Val
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Tyr	Ala	Lys	s Ala	Ser	· Asp	Leu	Tyr	Ile	Thr	Leu	Pro	Leu	Ala	Leu	Leu
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Phe	. Leu	ı Ile	e Val	L Arg	g Tyr	Phe	Phe	Glu	Leu	ı Tyr	· Val	Ala	Thr	Pro	Leu
	50)				55	5				60)			
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Asr	ı Ala	a Th	r Le	u Glı	u His	: Phe	е Туг	: Lev	ı Thi	r Sei	r Gly	/ Lys	s Gln	Pro	Lys

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Gln	Val	Glu	Val	Glu	Leu	Leu	Ser	Arg	Gln	Ser	Gly	Leu	Ser	Ģly	Arg
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Gln	Val	Glu	Arg	Trp	Phe	Arg	Arg	Arg	Arg	Asn	Gln	Asp	Arg	Pro	Ser
		115					120					125			
Leu	Leu	Lys	Lys	Phe	Arg	Glu	Ala	Ser	Trp	Arg	Phe	Thr	Phe	Tyr	Leu
	130					135					140				
Ile	Ala	Phe	Ile	Ala	Gly	Met	Ala	Val	Ile	Val	Asp	Lys	Pro	Trp	Phe
145					150					155					160
Tyr	Asp	Met	Lys	Lys	Val	Trp	Glu	Gly	Tyr	Pro	Ile	Gln	Ser	Thr	Ile
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Pro	Ser	Gln	Tyr	Trp	Tyr	Tyr	Met	Ile	Glu	Leu	Ser	Phe	Tyr	Trp	Ser
			180					185					190		
Leu	Leu	Phe	Ser	Ile	Ala	Ser	Asp	Val	Lys	Arg	Lys	Asp	Phe	Lys	Glu
		195					200					205			
Gln	Ile	Ile	His	His	Val	Ala	Thr	Ile	Ile	Leu	Ile	Ser	Phe	Ser	Trp
	210					215					220				
Phe	Ala	Asn	Tyr	Ile	Arg	Ala	Gly	Thr	Leu	Ile	Met	Ala	Leu	His	Asp
225	•		•		230					235					240
Ser	Ser	Asp	Tyr	Leu	Leu	Glu	Ser	Ala	Lys	Met	Phe	Asn	Tyr	Ala	Gly
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Trp	Lys	Asn	Thr	Cys	Asn	Asn	Ile	Phe	Ile	Val	Phe	Ala	Ile	Val	Phe
			260	ı				265	i				270		
Ile	Ile	Thr	Arg	Leu	Val	Ile	Leu	Pro	Phe	Trp	Ile	Leu	His	Cys	Thi
		275	;				280					285	;		

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Leu Val Tyr Pro Leu Glu Leu Tyr Pro Ala Phe Phe Gly Tyr Tyr Phe Phe Asn Ser Met Met Gly Val Leu Gln Leu Leu His Ile Phe Trp Ala Tyr Leu Ile Leu Arg Met Ala His Lys Phe Ile Thr Gly Lys Leu Val Glu Asp Glu Arg Ser Asp Arg Glu Glu Thr Glu Ser Ser Glu Gly Glu Glu Ala Ala Gly Gly Gly Ala Lys Ser Arg Pro Leu Ala Asn Gly His Pro Ile Leu Asn Asn Asn His Arg Lys Asn Asp <210> 5 <211> 585 <212> PRT <213> Homo sapiens <400> 5 Met Val Cys Arg Glu Gln Leu Ser Lys Asn Gln Val Lys Trp Val Phe Ala Gly Ile Thr Cys Val Ser Val Val Val Ile Ala Ala Ile Val Leu Ala Ile Thr Leu Arg Arg Pro Gly Cys Glu Leu Glu Ala Cys Ser Pro Asp Ala Asp Met Leu Asp Tyr Leu Leu Ser Leu Gly Gln Ile Ser Arg

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Lys	Asn	Ile	Lys	Ala	Val	Gly	Pro	Ser	Leu	Asp	Leu	Leų	Arg	Gln	Leu
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Thr	Glu	Glu	Gly	Lys	Val	Arg	Arg	Pro	Ile	Trp	Ile	Asn	Ala	Asp	Ile
				165					170					175	
Leu	Lys	Gly	Pro	Asn	Met	Leu	Ile	Ser	Thr	Glu	Val	Asn	Ala	Thr	Gln
			180					185					190		
Phe	Leu	Ala	Leu	Val	Gln	Glu	Lys	Tyr	Pro	Lys	Ala	Thr	Leu	Ser	Pro
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	210	•				215					220				
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Arg	Val	Thr	Phe	Pro	Val	Arg	Ser	Ser	Met	Val	Arg	Ala	Ala	Trp	Pro
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Trp	Gln	Ala	Ala	Ser	Asp	Pro	Met	Ser	Val	Glu	Asp	Leu	Leu	Tyr	Val
		275					280					285			
Arg	Asp	Asn	Thr	Ala	Val	His	Gln	Val	Tyr	Tyr	Asp	Ile	Phe	Glu	Pro
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Leu	Leu	Ser	Gln	Phe	Lys	Gln	Leu	Ala	Leu	Asn	Ala	Thr	Arg	Lys	Pro
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Met	Tyr	Tyr	Thr	Gly	Gly	Ser	Leu	Ile	Pro	Leu	Leu	Gln	Leu	Pro	Gly
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Asp	Asp	Gly	Leu	Asn	Val	Glu	Trp	Leu	Val	Pro	Asp	Val	Gln	Gly	Ser
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	370					375					380				
Val	His	Thr	Pro	Ser	G1 y	Asn	Ile	Leu	Thr	Leu	Glu	Ser	Cys	Leu	Gln
385					390					395					400
Gln	Leu	Ala	Thr	His	Pro	Gly	His	Trp.	Gly	Ile	His	Leu	G1n	Ile	Ala
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Glu	Pro	Ala	Ala	Leu	Arg	Pro	Ser	Leu	Ala	Leu	Leu	Ala	Arg	Leu	Ser
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Thr Ala Val Ala Glu Val Phe Pro His Val Thr Val Ala Pro Gly Trp Pro Glu Glu Val Leu Gly Ser Gly Tyr Arg Glu Gln Leu Leu Thr Asp Met Leu Glu Leu Cys Gln Gly Leu Trp Gln Pro Val Ser Phe Gln Met Gln Ala Met Leu Leu Gly His Ser Thr Ala Gly Ala Ile Gly Arg Leu Leu Ala Ser Ser Pro Arg Ala Thr Val Thr Val Glu His Asn Pro Ala Gly Gly Asp Tyr Ala Ser Val Arg Thr Ala Leu Leu Ala Ala Arg Ala Val Asp Arg Thr Arg Val Tyr Tyr Arg Leu Pro Gln Gly Tyr His Lys Asp Leu Leu Ala His Val Gly Arg Asn

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<212> PRT

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Pro	Val	Pro	Ser	Cys	Arg	Ala	Leu	Gln	Val	Leu	Lys	Pro	Arg	Asp	Arg
	50					55					60				
Ile	Ser	Ala	Ile	Ala	His	Arg	Gly	Gly	Ser	His	Asp	Ala	Pro	Glu	Asn
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Thr	Leu	Ala	Ala	Ile	Arg	Gln	Ala	Ala	Lys	Asn	Gly	Ala	Thr	Gly	Val
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Glu	Leu	Asp	Ile	Glu	Phe	Thr	Ser	Asp	Gly	Ile	Pro	Val	Leu	Met	His
			100					105					110	•	•
Asp	Asn	Thr	Val	Asp	Arg	Thr	Thr	Asp	Gly	Thr	Gly	Arg	Leu	Cys	Asp
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	130)				135					140				
Leu	Arg	, Asr	Asp	Phe	Pro	Asp	Glu	Lys	Ile	Pro	Thr	Leu	Arg	Glu	Ala
145	5				150)				155					160
Val	Ala	Glu	ı Cys	Leu	ı Asr	His	Asn	Leu	Ţhr	lle	Phe	Phe	Asp	Val	Lys
				165	5				170)				175	,
Gly	/ His	s Ala	a His	s Lys	s Ala	a Thr	Glu	ı Ala	. Lei	ı Lys	Lys	Me1	: Tyr	Met	Glu
			180)				185	5				190)	
Phe	e Pro	o Gla	n Lei	ı Tyı	r Ası	n Ası	ı Sei	r Val	l Val	l Cys	s Sea	r Phe	e Leu	Pro	Glu
		19	5 .				200)				20	5		
Vo	1 11.	a Tu	∽ 1 υ	e Ma	t Ar	~ G1:	n Thi	r Ası	n Ar	o Asi	o Va	1 11	e Thi	r Ala	a Leu

	210					215					220				
Thr	His	Arg	Pro	Trp	Ser	Leu	Ser	His	Thr	Gly	Asp	Gly	Lys	Pro	Are
225				,	230			•		235	•				240
Tyr	Asp	Thr	Phe	Trp	Lys	His	Phe	Ile	Phe	Val	Met	Met	Asp	Ile	Leu
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Leu	Asp	Trp	Ser	Met	His	Asn	Ile	Leu	Trp	Tyr	Leu	Cys	Gly	Ile	Sei
			260					265					270		
Ala	Phe	Leu	Met	G1n	Lys	Asp	Phe	Val	Ser	Pro	Ala	Tyr	Leu	Lys	Lys
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Trp	Ser	Ala	Lys	Gly	Ile	Gln	Val	Val	Gly	Trp	Thr	Val	Asn	Thr	Phe
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Asp	Glu	Lys	Ser	Tyr	Tyr	Glu	Ser	His	Leu	Gly	Ser	Ser	Tyr	Ile	Thr
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Glv	Ala	Ala	Ala	Val	Glv	Leu	Glv	Leu	Thr	Leu	Phe	Thr	Cys	Gly	Pro
,		35			,		40					45	•	•	
•					•					-1	ent.				71
His	Thr	Leu	His	Ser	Leu	Val	Thr	lle	Leu	Gly	Thr	Trp	Ala	Leu	IIe
	50					55					60				
Gln	Ala	Gln	Pro	Cys	Ser	Cys	His	Ala	Leu	Ala	Leu	Ala	Trp	Thr	Phe
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Ser	Tyr	Leu	Leu	Phe	Phe	Arg	Ala	Leu	Ser	Leu	Leu	Gly	Leu	Pro	Thr
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Pro	Thr	Pro	Phe	Thr	Asn	Ala	Val	Gln	Leu	Leu	Leu	Thr	Leu	Lys	Leu
			100					105					110		
Va1	Sor	Ī en		Ser	G1 ₁₁	Val	G1n		Leu	His	Leu	Ala	Gln	Arg	Lvs
101	GCI			501	014	,,,,	120					125			-,-
	,	115	_			_		41	_	~ 1	,		,	•	D
Glu	Met	Ala	Ser	Gly	Phe	Ser	Lys	Gly	Pro	Thr		Gly	Leu	Leu	Pro
	130					135					140				
Asp	Val	Pro	Ser	Leu	Met	Glu	Thr	Leu	Ser	Tyr	Ser	Tyr	Cys	Tyr	Val
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Gly	Ile	Met	Thr	Gly	Pro	Phe	Phe	Arg	Tyr	Arg	Thr	Tyr	Leu	Asp	Trp
				165					170					175	
Leu	Glu	Gln	Pro	Phe	Pro	Gly	Ala	Val	Pro	Ser	Leu	Arg	Pro	Leu	Leu
			180					185					190		
A ~~ ~	A	. 41.			41 a	Pro	וום ז		G1 v	וום ו	I en	Phe	Leu	l.en	Ser
wrR	vr.R			rio	VIG	110			UI,	Leu	Dea			200	
		195					200					205		_	
Ser	His	Leu	Phe	Pro	Leu	Glu	Ala	Val	Arg	Glu	Asp	Ala	Phe	Tyr	Ala
	210)				215					220)			
Arg	Pro	Leu	Pro	Ala	Arg	Leu	Phe	Tyr	Met	Ile	Pro	Val	Phe	Phe	Ala

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Phe Arg Met Arg Phe Tyr Val Ala Trp Ile Ala Ala Glu Cys Gly Cys Ile Ala Ala Gly Phe Gly Ala Tyr Pro Val Ala Ala Lys Ala Arg Ala Gly Gly Gly Pro Thr Leu Gln Cys Pro Pro Pro Ser Ser Pro Glu Lys Ala Ala Ser Leu Glu Tyr Asp Tyr Glu Thr Ile Arg Asn Ile Asp Cys Tyr Ser Thr Asp Phe Cys Val Arg Val Arg Asp Gly Met Arg Tyr Trp Asn Met Thr Val Gln Trp Trp Leu Ala Gln Tyr Ile Tyr Lys Ser Ala Pro Ala Arg Ser Tyr Val Leu Arg Leu <210> 8 **<211> 89** <212> PRT <213> Homo sapiens <400> 8 Met Tyr Met Gln Asp Tyr Trp Arg Thr Trp Leu Lys Gly Leu Arg Gly Phe Phe Phe Val Gly Val Leu Phe Ser Ala Val Ser Ile Ala Ala Phe

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Pro	Thr	Ser	Tyr	Tyr	Leu	Ser	Ser	Val	Trp	Ser	Phe	Ile	Ser	Phe	Lys
	50					55					60				
Trp	Ala	Phe	Leu	Leu	Ser	Leu	Tyr	Ala	His	Arg	Tyr	Arg	Ala	Asp	Phe
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Val	Met	Va1	Ala	Thr	Asn	Thr	Pro	Hic	Ser	Thr	Len	Ser	Val	Asn	Trn

				85					90					95	
Ser	Leu	Leu	Leu	Ser	Pro	Ģlu	Pro	Asp	Gly	Gly	Leu	Met	Va _. l	L _{eu}	Pro
		•	100					105				,	110		•
Lys	Asp	Ser	Ile	Gln	Phe	Ser	Ser	Ala	Leu	Val	Phe	Thr	Arg	Leu	Leu
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Glu	Phe	Asp	Ser	Thr	Asn	Val	Ser	Asp	Thr	Ala	Ala	Lys	Pro	Leu	Gly
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145					150					155					160
Thr	Asp	Ser	Leu	Asp	Pro	Ala	Thr	Leu	Ser	Ala	Thr	Phe	Gln	Gly	His
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Pro	Met	Asn	Asp	Pro	Thr	Arg	Thr	Phe	Ala	Asn	Gly	Ser	Leu _.	Ala	Phe
			180					185					190		
Arg	Val	Gln	Ala	Phe	Ser	Arg	Ser	Ser	Arg	Pro	Ala	Gln	Pro	Pro	Arg
		195					200					205			
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Asp	Asp	Glu	Tyr	Ala	Pro	Ala	Val	Phe	Gln	Leu	Asp	Gln	Leu	Leu	Trp
			260					265					270		
Gly	Ser	Leu	Pro	Ser	Gly	Phe	Ala	Gln	Trp	Arg	Pro	Val	Ala	Tyr	Ser
		275					280					285			

Gln	Lys	Pro	Gly	Gly	Arg	Glu	Ser	Ala	Leu	Pro	Cys	Gln	Ala	Ser	Pro
	290					295					300				٠
Leu	His	Pro	Ala	Leu	Ala	Tyr	Ser	Leu	Pro	Gln	Ser	Pro	Ile	Val	Arg
305					310					315					320
Ala	Phe	Phe	Gly	Ser	Gln	Asn	Asn	Phe	Cys	Ala	Phe	Asn	Leu	Thr	Phe
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Ser	Met	Leu	Leu	Gly	Val	Gly	Phe	Pro	Pro	Val	Asp	Gly	Leu	Ser	Pro
		355					360					365			
Leu	Val	Leu	Gly	Ile	Met	Ala	Val	Ala	Leu	Gly	Ala	Pro	Gly	Leu	Met
	370	•				375					380				
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Pro	Arg	Arg	Ser	Phe	Phe	Glu	Ser	Phe	Ile	Arg	Thr	Leu	Ile	Ile	Thr

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			20					25					30		
Cys	Val	Ala	Leu	Ala	Val	Val	Leu	Ser	Ser	Val	Ser	Ile	Cys	Asp	Gly
		35					40					45		÷	
His	Trp	Leu	Leu	Ala	Glu	Asp	Arg	Leu	Phe	Gly	Leu	Trp	His	Phe	Cys
	50					55					60				
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Ala	His	Val	Pro	Gly	Leu	Ala	Val	Gly	Met	Gly	Leu	Val	Arg	Ser	Val
				85					90					95	
Gly	Ala	Leu	Ala	Val	Val	Ala	Ala	Ile	Phe	G1y	Leu	Glu	Phe	Leu	Met
			100					105					110		
Val	Ser	Gln	Leu	Cys	Glu	Asp	Lys	His	Ser	Gln	Cys	Lys	Trp	Val	Met
		115					120				•	125			
Gly	Ser	Ile	Leu	Leu	Leu	Val	Ser	Phe	Val	Leu	Ser	Ser	Gly	Gly	Leu
	130					135					140				
Leu	Gly	Phe	Val	Ile	Leu	Leu	Arg	Asn	Gln	Val	Thr	Leu	Ile	Gly	Phe
145					150					155					160
Thr	Leu	Met	Phe	Trp	Cys	Glu	Phe	Thr	Ala	Ser	Phe	Leu	Leu	Phe	Leu
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<210> 11

<211> 801

<212> DNA

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7017		
V Z I J Z	rithiu	Saurens
\U	***	sapiens

⟨400⟩ 11

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<211> 1257

<212> DNA

<213> Homo sapiens

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gtgaeggtge aggageeegg eegeeggeee eegeteaegt ttegeatega eegeggegee 180

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240	gctgccctc	tgctggcgcc	cgcggccagg	ggccgaggtc	actccccaa	tacgggcttg
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360	gaaaatatca	cgtttaaaga	ggaaactgca	gctgcagagg	ggattgcctt	atcaaacagt
420	caaagaggag	ataataaatc	gtcatctaca	agttgctgta	tccacaatgc	cgggccgctt
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600	cgtgtcaata	ctctagtctt	agccgtggct	gaagaacttc	gaatgccacc	gttggaactc
660	cattcagaag	tattctactt	gcatggctca	tatttcttca	ttttgatgat	tcctttattg
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1140	aatcaacatt	gaacaggaga	ctcactccga	ggatggggag	ctcttcctca	gggatctcac
1200	cctcacactc	tcctcagtgc	agttttggcc	tattattgcc	aagaatggtt	gcagtaacaa
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⟨210⟩ 13

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<212> DNA

<213> Homo sapiens

<400> 13

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240	catagacctc	atttccgatt	gtcgttctca	aggaaaagta	cagttcccga	tggaaaatca
300	tgccaatggc	acaatggcca	gtggatgtgt	ctatgacttt	acctgtgccg	gagagtgaca
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480	ccttgacaga	gtggaggact	gatcagtatt	cgaaagaggg	ctgaaccaaa	ttctccgctg
540	aggagtcact	attaccctgc	ccagaccggg	ccccaactgg	cttttaaaac	ccttccggct
600	tgagaagttt	aattaaagtt	cagcttatag	cccaaagaat	acattgtagc	tgtgtgtggc
660	tggcggggaa	ctgtgtttaa	gattatgtgg	ctgccgatat	gagataacta	gatgtggagc
720	gccaattgtg	gtccacctgc	tgtggtgata	tggaaagtat	ctagaagaat	gtcaacgatg
780	tgcagatggg	taagtttaac	ttatcagact	tattcagttt	atgaacttct	tctgagagaa
840	acagcctgtc	caactacaga	aaactgccta	caggccaaaa	actacatatt	tttattggtc
900	tcaacaaaag	tggccttgtg	aaaaccaccg	cacgggttta	tccctgtaac	accaccacat
960	attagccggc	gtgactttgt	tattgttcaa	ggagggcaat	cggggactct	tgtagacgga
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1080	tgccaggctg	agaacatgag	caggcgggca	ggcgattcag	agggaaattt	atctacaaag
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1200	gatgttcaag	gctttatcat	atgccaaaca	aggcaaaatc	aagatgggcg	caagtaggtg
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⟨210⟩ 14

⟨211⟩ 1140

<212> DNA

<213> Homo sapiens

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<400> 14

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180	gctgtacgtg	acttctttga	atcgttcgat	gctcttcctc	ccctggcctt	atcacgctgc
240	ggcacctccc	ctcggctgcg	aaggagaaaa	cttgaacata	tggctgccct	gctacaccac
300	ggtggaagta	agcccaagca	agtggcaagc	ctacctgacc	tggaacattt	aacgccacct
360	gttccgtcgc	tagagcgttg	ggccgccagg	cgggctctct	cccggcagag	gagċttttgt
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480	accctggttc	ttgtggataa	atggccgtca	cattgccggc	tgattgcctt	acattttacc
540	ttcccagtat	gcactatccc	cccatacaga	ggagggatat	agaaagtttg	tatgacatga
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660	cattctcatc	tggccaccat	atccaccatg	ggaacagatc	aggatttcaa	gtcaagcgaa
720	tctgcatgac	taatcatggc	gctgggactc	ttacatccga	ggtttgccaa	agcttttcct
780	gaagaacacc	acgcgggatg	atgtttaact	gtcagccaag	acctgctgga	tcttccgatt
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900	tgccttcttt	agctctatcc	tacccactgg	caccctggtg	tcctgcattg	cccttctgga
960	cttctgggcc	tgctgcatat	gttctacagc	catgatggga	tcttcaattc	ggctattact
1020	agatgaacgc	agctggtaga	ataactggaa	ccacaagttc	tgcgcatggc	tacctcattt
1080	gggaggagca	ctgcagctgg	ggggaggagg	gagctcagag	aagaaacaga	agtgaccggg
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⟨210⟩ 15

⟨211⟩ 1755

<212> DNA

<213> Homo sapiens

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⟨400⟩ 15

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180	gagcctgggc	actacctgct	gacatgctgg	ccctgatgcc	aggcctgcag	tgtgagctgg
240	cagcaagaaa	acgcagccaa	acctggtacc	cttggaggtc	ggcgagatgc	cagatcagcc
300	caatgtagaa	aggctgacgt	acagtcctgg	cagcaacatc	ctgccctgaa	gccatgacag
360	cactatctac	cacacccccc	cccatcatgg	gacaggagtt	cagccaatga	gggctcggca
420	aaagggcatc	gctcttccca	gctgtgctgg	gtggctggac	cactggagca	agtgacaaca
480	gcggcagctg	tggacctcct	ggccctccc	caaggcagtg	tcaagaacat	aaactggact
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600	ccaggagaag	tggccctggt	acacagttcc	ggtcaatgcc	tctcaactga	aacatgctca
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720	agtgccccag	tggtgggagg	atgcacgagc	ggtggagaag	cccaagccat	aggacgtaca
780	cttcagctgg	cctggcccca	gtgcgggctg	gtcttccatg	tccctgtacg	agggtcacct
840	ggaccccatg	aggctgcctc	acgctgtggc	gtacagcctg	aatctgagag	ctgctgagcc
. 900	ctactatgac	tccaccaagt	aacactgctg	cgtccgggat	atctgctcta	tcggtggaag
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⟨211⟩ 1035

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<213> Homo sapiens

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Met Val Lys Ile Ser Phe Gln	
1 5	
ccc gcc gtg gct ggc atc aag ggc gac aag gct gac aag gcg tcg gcg	159
Pro Ala Val Ala Gly Ile Lys Gly Asp Lys Ala Asp Lys Ala Ser Ala	
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Ser Ala Pro Ala Pro Ala Ser Ala Thr Glu Ile Leu Leu Thr Pro Ala

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Met Asn Val Lys Arg Gly Thr Tyr	Leu Pro Gln T	hr Tyr Ile Ile Gln	
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gag gag atg gtg gtc acg gag cat	gtc agt gac as	ag gag gcc ctg ggg	735
Glu Glu Met Val Val Thr Glu His	Val Ser Asp L	ys Glu Ala Leu Gly	
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Ser Phe Ile Tyr His Leu Cys Asn	Gly Lys Asp Ti	hr Tyr Arg Leu Arg	
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Arg Arg Ala Thr Arg Arg Arg Ile	Asn Lys Arg G	ly Ala Lys Asn Cys	
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Asn Ala Ile Arg His Phe Glu Asn	Thr Phe Val Va	al Glu Thr Leu Ile	
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Cys Gly Val Val			
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Lys	Lys	Ala	Ile	Ser	Lys	Leu	Thr	Thr	Arg	Thr	Val	Lys	Lys	Gly	Asp	
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Glu Val Glu Trp Pho	е		· ·	
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		Met Arg Gly	Ala Asn Ala Trp	Ala
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cca ctc tgc ctg c	tg ctg gct gcc	gcc acc cag	ctc tcg cgg cag	cag 281
Pro Leu Cys Leu L	eu Leu Ala Ala	Ala Thr Gln	Leu Ser Arg Gln	Gln
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Ser	Gly	Phe	Ile	Gly	Ser	Glu	Gly	Phe	Pro	Gly	Val	Tyr	Pro	Pro	Asn	
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Ser	Lys	Cys	Thr	Trp	Lys	Ile	Thr	Val	Pro	Glu	Gly	Lys	Val	Val	Val	
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Leu	Asn	Phe	Arg	Phe	Ile	Asp	Leu	Glu	Ser	Asp	Asn	Leu	Cys	Are	Tyr	
		75				٠	80					85	,			
gac	ttt	gtg	gat	gtg	tac	aat	ggc	cat	gcc	aat	ggc	cag	g cgc	ati	ggc	521
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Arg	Phe	Cys	Gly	Thr	Phe	Arg	Pro	Gly	Ala	Leu	ı Val	Se	r Se	r Gl	y Asn	ı
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Lys	Me	t Me	t Val	l Glr	n Met	: Ile	Sei	r Ası	Ala	a Ası	n Thi	r Al	a Gl	y As	n Gly	<i>i</i>
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Phe	e Me	t Al	a Me	t Ph	e Sei	r Ala	a Ala	a Gl	u Pro	o Asi	n Gl	u Ar	g Gl	y As	p Gli	n
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+01	+ + ~	+ ~~	o 00	a ct	c cti	t ga	c ag	асс	t tc	C gg	c tc	t tt	t as	a ac	c cc	c 713

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Asn	Trp	Pro	Asp	Arg	Asp	Tyr	Pro	Ala	Gly	Val	Thr	Cys	Val	Trp	His	
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Ile	Val	Ala	Pro	Lys	Asn	Gln	Leu	Ile	Glu	Leu	Lys	Phe	Glu	Lys	Phe	
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Asp	Val	Glu	Arg	Asp	Asn	Tyr	Cys	Arg	Tyr	Asp	Tyr	Val	Ala	Val	Phe	
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Asp	Ser	Pro	Pro	Ala	Pro	Ile	Val	Ser	G1 ụ	Arg	Asn	Glu	Leu	Leu	Ile	
		235					240					245				
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Gln	Phe	Leu	Ser	Asp	Leu	Ser	Leu	Thr	Ala	Asp	Gly	Phe	Ile	Gly	His	
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Tyr	Ile	Phe	Arg	Pro	Lys	Lys	Leu	Pro	Thr	Thr	Thr	Glu	Gln	Pro	Val	
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Thr	Thr	Thr	Phe	Pro	Val	Thr	Thr	Gly	Leu	Lys	Thr	Thr	Val	Ala	Leu	

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(Cys	Gln	Gln	Lys	Cys	Arg	Arg	Thr	Gly	Thr	Leu	Glu	Gly	Asn	Tyr	Cys	
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1	tca	agt	gac	ttt	gta	tta	gcc	ggc	act	gtt	atc	aca	acc	atc	act	cgc	1193
5	Ser	Ser	Asp	Phe	Val	Leu	Ala	Gly	Thr	Val	Ile	Thr	Thr	Ile	Thr	Arg	
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	Ile	Ile	Met	Gly	Gln	Val	Gly	Glu	Asp	Gly	Arg	Gly	Lys	Ile	Met	Pro	
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	Asn	Ser	Phe	Ile	Met	Met	Phe	Lys	Thr	Lys	Asn	Gln	Lys	Leu	Leu	Asp	
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	gcc	tta	888	aat	aag	caa	tgt	taa	cagt	gaa (ctgt	gtcc	at t	taag	С		1480
	Ala	Leu	Lys	Asn	Lys	Gln	Cys										
		410					415										

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Met Leu Gln

1

acc ttg tat gat tac ttc tgg tgg gaa cgt ctg tgg ctg cct gtg aac 281

Thr Leu Tyr Asp Tyr Phe Trp Trp Glu Arg Leu Trp Leu Pro Val Asn

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Leu	Thr	Trp	Ala	Asp	Leu	Glu	Asp	Arg	Asp	Gly	Arg	Val	Tyr	Ala	Lys	
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gcc	tca	gat	ctc	tat	atc	acg	ctg	ccc	ctg	gcc	ttg	ctc	ttc	ctc	atc	377
Ala	Ser	Asp	Leu	Tyr	Ile	Thr	Leu	Pro	Leu	Ala	Leu	Leu	Phe	Leu	Ile	
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Val	Arg	Tyr	Phe	Phe	Glu	Leu	Tyr	Val	Ala	Thr	Pro	Leu	Ala	Ala	Leu	
			55					60					65			
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Leu	Asn	Ile	Lys	Glu	Lys	Thr	Arg	Leu	Arg	Ala	Pro	Pro	Asn	Ala	Thr	
		70					75				•	80				
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Leu	Glu	His	Phe	Tyr	Leu	Thr	Ser	Gly	Lys	Gln	Pro	Lys	Gln	Val	Glu	
	85					90					95					
gta	gag	ctt	ttg	tcc	cgg	cag	agc	ggg	ctc	tct	ggc	cgc	cag	gta	gag	569
Val	Glu	Leu	Leu	Ser	Arg	Gln	Ser	Gly	Leu	Ser	Gly	Arg	Gln	Val	Glu	
100					105					110					115	
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Arg	Trp	Phe	Arg	Arg	Arg	Arg	Asn	Gln	Asp	Arg	Pro	Ser	Leu	Leu	Lys	
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Lys	Phe	Arg	Glu	Ala	Ser	Trp	Arg	Phe	Thr	Phe	Tyr	Leu	Ile	Ala	Phe	
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Lys	Lys	Val	Trp	Glu	Gly	Tyr	Pro	Ile	Gln	Ser	Thr	Ile	Pro	Ser	Gln	
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Tyr	Trp	Tyr	Tyr	Met	Ile	Glu	Leu	Ser	Phe	Tyr	Trp	Ser	Leu	Leu	Phe	
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Ser	Ile	Ala	Ser	Asp	Val	Lys	Arg	Lys	Asp	Phe	Lys	Glu	Gln	Ile	Ile	
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His	His	Val	Ala	Thr	Ile	Ile	Leu	Ile	Ser	Phe	Ser	Trp	Phe	Ala	Asn	
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Tyr	Ile	Arg	Ala	Gly	Thr	Leu	Ile	Met	Ala	Leu	His	Asp	Ser	Ser	Asp	
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Tyr	Leu	ı Let	ı Glu	Ser	Ala	Lys	Met	. Phe	: Asr	1 Tyr	· Ala	Gly	Tr	Lys	s Asn	
	245	5				250)				25	5				
aco	tgo	88	e aac	ato	tto	ato	gto	tto	gco	att	gt	t tti	ato	ato	acc	1049
Thi	r Cys	s Ası	n Asr	ı Ile	Phe	e Ile	e Val	l Phe	e Ala	a Ile	e Va	l Phe	e Ile	e Il	e Thr	
260)				265	5				270)				275	
cg	a ct	g gt	c ato	ctg	g cc	tto	tg	g ato	ct	g cat	t tg	C 8C	c ct	g gt	g tac	1097

Arg l	Leu	Val	Ile	Leu	Pro	Phe	Trp	Ile	Leu	His	Cys	Thr	Leu	Val	Tyr	
. •				280					285					290		
cca (ctg	gag	ctc-	tat	cct	gcc	ttc	ttt	ggc	tat	tac	ttc	ttc	aat	tcc	1145
Pro :	Leu	Glu	Leu	Tyr	Pro	Ala	Phe	Phe	Gly	Tyr	Tyr	Phe	Phe	Asn	Ser	
			295					300					305			
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Met	Met	Gly	Val	Leu	Gln	Leu	Leu	His	Ile	Phe	Trp	Ala	Tyr	Leu	Ile	
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Leu	Arg	Met	Ala	His	Lys	Phe	Ile	Thr	Gly	Lys	Leu	Val	Glu	Asp	Glu	
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Arg	Ser	Asp	Arg	Glu	Glu	Thr	Glu	Ser	Ser	Glu	Gly	Glu	ı Glu	Ala	Ala	
340					345	j				350)				355	
gct	ggg	gga	gga	gca	aag	ago	cgg	cco	cta	a gcc	aat	gge	cac	ccc	atc	1337
Ala	Gly	Gly	Gly	Ala	a Lys	Ser	Arg	Pro	Let	ı Ala	A Asr	G1;	y His	Pro	Ile	
				360)				36	5				370)	
ctc	aat	t aad	c aac	c ca	t cg1	t aag	g aat	ga.	c tg	aacca	atta	ttc	cagc	tgc (tccca	1390
Leu	Ası	n Ası	n Ası	n His	s Ar	g Ly:	s Asr	n As	р .							
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<213> Homo sapiens

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<221> CDS

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agcgtcgcc atg gtc tgc agg gag cag tta tca aag aat cag gtc aag 168
Met Val Cys Arg Glu Gln Leu Ser Lys Asn Gln Val Lys

1 5 10

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Ile	Val	Leu	Ala	Ile	Thr	Leu	Arg	Arg	Pro	Gly	Cys	Glu	Leu	Glu	Ala	
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tgc	agc	cct	gat	gcc	gac	atg	ctg	gac	tac	ctg	ctg	agc	ctg	ggc	cag	312
Cys	Ser	Pro	Asp	Ala	Asp	Met	Leu	Asp	Tyr	Leu	Leu	Ser	Leu	Gly	Gln	
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atc	agc	cgg	cga	gat	gcc	ttg	gag	gtc	acc	tgg	tac	cac	gca	gcc	aac	360
Ile	Ser	Arg	Arg	Asp	Ala	Leu	Glu	Val	Thr	Trp	Tyr	His		Ala	Asn	
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				atg												408
Ser	Lys		Ala	Met	Thr	Ala		Leu	Asn	Ser	Asn		Thr	Val	Leu	
		80					85					90				450
		_		aat												456
Glu		Asp	Val	Asn	Val		GLY	Leu	Gly	Ihr		Asn	Glu	Inr	GIY	
	95					100			_ 4 _		105				-4-	504
				gca								-				504
	Pro	116	мет	Ala		Pro	rro	inr	116		5er	ASP	ASN	ınr	125	
110		***	.+-	~~~	115	ata	ota	~~~	t at	120		222	~~~	ata		552
				gac Asp												332
OIU	OIII	rrb	ren	130	VIQ	191	Leu	GIÀ	135	Jer.	OIII	Lys	GIÀ	140	LJS	
ctø	gan	t +c	220	aac	atr	ลลฮ	8 08	gto		ccc	tee	ctø	gar		ctø	600
	_		_	Acn												

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Arg	Gln	Leu	Thr	Glu	Glu	Gly	Lys	Val	Arg	Arg	Pro	Ile	Trp	Ile	Asn	
		160					165					170				
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Ala	Asp	Ile	Leu	Lys	Gly	Pro	Asn	Met	Leu	Ile	Ser	Thr	Glu	Val	Asn	
	175					180					185					
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Ala	Thr	Gln	Phe	Leu	Ala	Leu	Va1	Gln	Glu	Lys	Tyr	Pro	Lys	Ala	Thr	
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cta	tct	cca	ggc	tgg	acc	acc	ttc	tac	atg	tcc	acg	tcc	cca	aac	agg	792
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Thr	Tyr	Thr	Gln	Ala	Met	Val	G1u	Lys	Met	His	G1u	Leu	Val	Gly	Gly	
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Val	Pro	Gln	Arg	Val	Thr	Phe	Pro	Val	Arg	Ser	Ser	Met	Val	Arg	Ala	
		240				•	245	٠				250				
gcc	tgg	ccc	cac	ttc	agc	tgg	ctg	ctg	agc	caa	tct	gag	agg	tac	agc	936
Ala	Trp	Pro	His	Phe	Ser	Trp	Leu	Leu	Ser	Gln	Ser	Glu	Arg	Tyr	Ser	
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Leu	Tyŗ	Val	Arg	Asp	Asn	Thr	Ala	Val	His	Gln _.	Val	Tyr	Tyr	Asp	Ile	
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Phe	Glu	Pro	Leu	Leu	Ser	Gln	Phe	Lys	Gln	Leu	Ala	Leu	Asn	Ala	Thr	
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Leu	Pro	Gly	Asp	Asp	Gly	Leu	Asn	Val	Glu	Trp	Leu	Val	Pro	Asp	Val	
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		400		•			405					410				
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Gln	lle	Ala	Glu	Pro	Ala	Ala	Leu	Arg	Pro	Ser	Leu	Ala	Leu	Leu	Ala	
	415					420					425					•
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Arg	Leu	Ser	Ser	Leu	Gly	Leu	Ļeu	His	Trp	Pro	Val	Trp	Val	Gly	Ala	
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888	atc	tcc	cac	ggg	agt	ttt	tcg	gtc	ccc	ggc	cat	gtg	gct	ggc	aga	1512
Lys	Ile	Ser	His	Gly	Ser	Phe	Ser	Val	Pro	Gly	His	Val	Ala	Gly	Arg	
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Glu	Leu	Leu	Thr	Ala	Val	Ala	Glu	Val	Phe	Pro	His	Val	Thr	Val	Ala	
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Pro	Gly	Trp	Pro	Glu	Glu	Val	Leu	Gly	Ser	Gly	Tyr	Arg	Glu	Gln	Leu	
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Leu	Thr	Asp	Met	Leu	Glu	Leu	Cys	Gln	Gly	Leu	Trp	Gln	Pro	Val	Ser	
	495					500					505					
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545 550 555 1848 get agg get gtg gac agg acc cga gtc tac tac agg cta ccc cag ggc Ala Arg Ala Val Asp Arg Thr Arg Val Tyr Tyr Arg Leu Pro Gln Gly 560 565 570 tac cac aag gac ttg ctg gct cat gtt ggt aga aac tgagcaccca ggggtg 1900 Tyr His Lys Asp Leu Leu Ala His Val Gly Arg Asn 575 580 585 1960 gtgggccagc ggacctcagg gcggaggctt cccacgggga ggcaggaaga aataaaggtc tttggctttc tcc 1973 <210> 26 <211> 1606 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (135)...(1130) <400> 26 attgtgcggc gctggtcccc tcagagggtt cctgctgctg ccggtgcctt ggaccctccc 60 cctcgcttct cgttctactg ccccaggagc ccggcgggtc cgggactccc gtccgtgccg 120 gtgcggcgc cggc atg tgg ctg tgg gag gac cag ggc ggc ctc ctg ggc 170 Met Trp Leu Trp Glu Asp Gln Gly Gly Leu Leu Gly 1 5 10 218 cct ttc tcc ttc ctg ctg cta gtg ctg ctg ctg gtg acg cgg agc ccg Pro Phe Ser Phe Leu Leu Leu Val Leu Leu Val Thr Arg Ser Pro

		15					20					25				
gtc	aat	gcc	tgc	ctc	ctc	acc	ggc	agc	ctc	ttc	gtt	cta	ctg	cgc	gtc	 266
Val	Asn	Ala	Cys	Leu	Leu	Thr	Gly	Ser	Leu	Phe	Val	Leu	Leu	Arg	Val	
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ttc	agc	ttt	gag	ccg	gtg	ССС	tct	tgc	agg	gcc	ctg	cag	gtg	ctc	aag	314
Phe	Ser	Phe	Glu	Pro	Val	Pro	Ser	Cys	Arg	Ala	Leu	Gln	Val	Leu	Lys	
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ccc	cgg	gac	cgc	att	tct	gcc	atc	gcc	cac	cgt	ggc	ggc	agc	cac	gac	362
Pro	Arg	Asp	Arg	Ile	Ser	Ala	Ile	Ala	His	Arg	Gly	Gly	Ser	His	Asp	
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gcg	ccc	gag	aac	acg	ctg	gcg	gcc	att	cgg	cag	gca	gct	aag	aat	gga	410
Ala	Pro	Glu	Asn	Thr	Leu	Ala	Ala	Iļe	Arg	Gln	Ala	Ala	Lys	Asn	Gly	
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Ala	Thr	Gly	Val	Glu	Leu	Asp	Ile	Glu	Phe	Thr	Ser	Asp	Gly	Ile	Pro	
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gtc	tta	atg	cac	gat	aac	aca	gta	gat	agg	acg	act	gat	ggg	act	ggg	506
Val	Leu	Met	His	Asp	Asn	Thr	Val	Asp	Arg	Thr	Thr	Asp	Gly	Thr	Gly	
	110					115					120			•		
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Arg	Leu	Cys	Asp	Leu	Thr	Phe	Glu	Gln	Ile	Arg	Lys	Leu	Asn	Pro	Ala	
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gca	aac	cac	aga	ctc	agg	aat	gat	ttc	cct	gat	gaa	aag	atc	cct	acc	602
Ala	Asn	His	Arg	Leu	Arg	Asn	Asp	Phe	Pro	Asp	G1u	Lys	Ile	Pro	Thr	
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cta	agg	gaa	gct	gtt	gca	gag	tgc	cta	aac	cat	aac	ctc	aca	atc	ttc	650
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			160					165					170		•	
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Phe	Asp	Val	Lys	Gly	His	Ala	His	Lys	Ala	Thr	Glu	Ala	Leu	Lys	Lys	
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Met	Tyr	Met	Glu	Phe	Pro	Gln	Leu	Tyr	Asn	Asn	Ser	Val	Val	Cys	Ser	
	190					195					200					
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Phe	Leu	Pro	Glu	Val	Ile	Tyr	Lys	Met	Arg	Gln	Thr	Asp	Arg	Asp	Val	
205					210					215		•			220	
ata	aca	gca	tta	act	cac	aga	cct	tgg	agc	cta	agc	cat	aca	gga	gat	842
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Met	Asp	Ile	Leu	Leu	Asp	Trp	Ser	Met	His	Asn	Ile	Lev	Trp	Туг	Leu	
		255	5				260)				265	5			
tgt	gga	att	tca:	gct	tto	cto	ate	caa	aag	gat	ttt	gta	tce	င ငင္ရ	g gcc	986
Cys	Gly	ı Ile	e Ser	Ala	. Phe	Leu	ı Met	: Gln	Lys	: Asp	Phe	· Val	l Sei	r Pro	Ala	
	270)				275	5				280)				
tac	ttg	g aag	g aag	g tgg	g tca	a gct	. aaa	a gga	ato	ca;	g gti	t gti	t gg	t tg	g act	1034

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Tyr Leu Lys Lys Trp Ser Ala Lys Gly Ile Gln Val Val Gly Trp Thr	
285 290 295 300	
gtt aat acc ttt gat gaa aag agt tac tac gaa tcc cat ctt ggt tcc	1082
Val Asn Thr Phe Asp Glu Lys Ser Tyr Tyr Glu Ser His Leu Gly Ser	
305 310 315	
age tat ate act gae age atg gta gaa gae tge gaa eet cae tte	1127
Ser Tyr Ile Thr Asp Ser Met Val Glu Asp Cys Glu Pro His Phe	
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tag actttcacgg tgggacgaaa cgggttcaga aactgccagg ggcctcatac	1180
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taaagg	1606

⟨210⟩ 27

⟨211⟩ 2380

<212> DNA

<213> Homo sapiens

⟨220⟩

<221> CDS

〈222〉 (247)...(1284)

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ceggegggeg ce	cagaagga gcaggcgg	cg cggggggggg	ctgggcgggg gaggcgt	:ggc 180
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cagacc atg tc	g cct gaa gaa tgg	acg tat cta g	tg gtt ctt ctt ato	288
Met Se	r Pro Glu Glu Trp	Thr Tyr Leu V	al Val Leu Leu Ile	•
1	5		10	
tcc atc ccc a	tc ggc ttc ctc tt	t aag aaa gcc	ggt cct ggg ctg as	ag 336
Ser Ile Pro I	le Gly Phe Leu Ph	e Lys Lys Ala	Gly Pro Gly Leu Ly	ys
15	20	25		30
aga tgg gga g	ca gcc gct gtg gg	c ctg ggg ctc	acc ctg ttc acc t	gt <u>384</u>
Arg Trp Gly A	ala Ala Ala Val Gl	y Leu Gly Leu	Thr Leu Phe Thr C	ys
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Gly Pro His T	Thr Leu His Ser Le	u Val Thr Ile	Leu Gly Thr Trp A	la
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Leu Ile Gln A	Ala Gln Pro Cys Se	r Cys His Ala	Leu Ala Leu Ala T	rp
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Thr Phe Ser 1	Tyr Leu Leu Phe Ph	e Arg Ala Leu	Ser Leu Leu Gly L	eu
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Pro Thr Pro	Thr Pro Phe Thr As	n Ala Val Gln	Leu Leu Leu Thr L	eu

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agg	aag	gaa	atg	gcc	tca	ggc	ttc	agc	aag	ggg	ссс	acc	ctg	ggg	ctg	672
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Asp	Trp	Leu	Glu	Gln	Pro	Phe	Pro	Gly	Ala	Val	Pro	Ser	Leu	ı Arg	Pro	
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Leu	Leu	Arg	, Arg	Ala	Trp	Pro	Ala	Pro	Leu	Phe	Gly	Leu	Let	ı Phe	e Leu	
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Туз	- Ala	a Arı	g Pro	Lei	ı Pro	Ala	Arg	g Leu	Phe	Ty	r Me	t Ile	Pr	o Va	1 Phe	
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Phe _.	Ala	Phe	Arg	Met	Arg	Phe	Tyr	Val	Ala	Trp	Ile	Ala	Ala	Glu	Cys	
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Tyr	Trp	Asn	Met	Thr	Val	Gln	Trp	Trp	Leu	Ala	Gln	Tyr	Ile	Tyr	Lys	
	320					325					330					
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Ser	Ala	Pro	Ala	Arg	Ser	Tyr	Val	Leu	Arg	Leu	l					
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<210> 28

<211> 2017

<212> DNA

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<220>

<221> CDS

⟨222⟩ (360)...(629)

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<212> DNA

<213> Homo sapiens

PCT/JP00/05356

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His	s Se	r Thi	r Lei	ı Sei	r Val	Asn	Trp	Ser	Leu	ı Le	u Lei	u Sei	r Pr	o Gl	u Pro	•
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<212> DNA

<213> Homo sapiens

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<221> CDS

⟨222⟩ (53)...(631)

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Arg	Arg	Ser	Phe	Phe	Glu	Ser	Phe	Ile	Arg	Thr	Leu	Ile	Ile	Thr	Су	s	
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Val	Ala	Leu	Ala	Val	Val	Leu	Ser	Ser	Val	Ser	Ile	Cys	Asp	Gly	Hi	.s	
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Trp	Leu	ı Lev	ı Ala	Glu	Asp	Arg	Leu	Phe	Gly	Leu	Trp	His	Phe	Cy:	s Th	ır	
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acc	aco	c aac	c cag	gagt	gtg	ccg	ato	tgo	ttc	aga	gac	cte	gg	ca	g go	CC	295 ⁻
Thi	Thi	r Ası	n Glı	Ser	Val	Pro	Ile	Cys	Phe	Arg	Asp	Leu	ı Gly	y Gla	n A	la	
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Ala	Leu	Ala	Tyr	Gly	Ser	Leu	Leu	Leu	Met	Ala	Leu	Leu	Pro	Ile	Phe		
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Phe	Gly	Ala	Leu	Arg	Ser	Val	Arg	Cys	Ala	Arg	Gly	Lys	Asn	Ala	Ser		
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Asp	Met	Pro	Glu	Thr	Ile	Thr	Ser	Arg	Asp	Ala	Ala	Arg	Phe	Pro	Ile		
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Ile	Ala	Ser	Cys	Thr	Leu	Leu	Gly	Leu	Tyr	Leu	Phe	Phe	Lys	Ile	Phe		
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105

Gly Ile Leu Ala Leu Ser His Thr Ile Ser Pro Phe Met Asn Lys Phe $\,$

110

100

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Lys	Leu	Val	Phe	Pro	Gln	Asp	Leu	Leu	Glu	Lys	Gly	Leu	Glu	Ala	Asn
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Asn	Phe	Ala	Met	Leu	Gly	Leu	Gly	Asp	Val	Val	Ile	Pro	Gly	Ile	Phe
			260					265	:				270		
Ile	Ala	Leu	Leu	Leu	Arg	Phe	Asp	Ile	Ser	Leu	Lys	Lys	Asn	Thr	His
		275					280					285			
Thr	Tyr	Phe	Tyr	Thr	Ser	Phe	Ala	Ala	Tyr	Ile	Phe	Gly	Leu	Gly	Leu
	290					295					300				
Thr	Ile	Phe	Ile	Met	His	Ile	Phe	Lys	His	Ala	Gln	Pro	Ala	Leu	Leu
305					310					315					320

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Tyr Leu Val Pro Ala Cys Ile Gly Phe Pro Val Leu Val Ala Leu Ala Lys Gly Glu Val Thr Glu Met Phe Ser Tyr Glu Glu Ser Asn Pro Lys Asp Pro Ala Ala Val Thr Glu Ser Lys Glu Gly Thr Glu Ala Ser Ala Ser Lys Gly Leu Glu Lys Lys Glu Lys <210> 32 ⟨211⟩ 81 <212> PRT <213> Homo sapiens <400> 32 Met Thr Ala His Ser Phe Ala Leu Pro Val Ile Ile Phe Thr Thr Phe Trp Gly Leu Val Gly Ile Ala Gly Pro Trp Phe Val Pro Lys Gly Pro Asn Arg Gly Val Ile Ile Thr Met Leu Val Ala Thr Ala Val Cys Cys Tyr Leu Phe Trp Leu Ile Ala Ile Leu Ala Gln Leu Asn Pro Leu Phe Gly Pro Gln Leu Lys Asn Glu Thr Ile Trp Tyr Val Arg Phe Leu Trp Glu

Lys	Pro	Pro	Gln	Ile	Val	Val	Lys	Cys	Leu	Ala	Ala	Ala	Ala	Ile	Leu
145					150					155					160
Phe	Ile	Ser	Thr	Val	Asn	Ser	Leu	Ser	Val	Arg	Leu	Gly	Ser	Tyr	Val
				165					170					175	
Gln	Asn	Ile	Phe	Thr	Ala	Ala	Lys	Leu	Val	Ile	Val	Ala	Ile	Ile	Ile
			180					185					190		
Ile	Ser	Gly	Leu	Val	Leu	Leu	Ala	Gln	Gly	Asn	Thr	Lys	Asn	Phe	Asp
		195					200					205			
Asn	Ser	Phe	Glu	Gly	Ala	Gln	Leu	Ser	Val	Gly	Ala	Ile	Ser	Leu	Ala
	210					215					220				
Phe	Tyr	Asn	Gly	Leu	Trp	Ala	Tyr	Asp	Gly	Trp	Asn	Gln	Leu	Asn	Tyr
225					230					235					240
Ile	Thr	Glu	Glu	Leu	Arg	Asn	Pro	Tyr	Arg	Asn	Leu	Pro	Leu	Ala	Ile
				245					250					255	
Ile	Ile	Gly	Ile	Pro	Leu	Val	Thr	Ala	Cys	Tyr	Ile	Leu	Met	Asn	Val
			260					265					270		
Ser	Tyr	Phe	Thr	Val	Met	Thr	Ala	Thr	Glu	Leu	Leu	Gln	Ser	Gln	Ala
		275					280					285			
Val	Ala	Val	Thr	Phe	Gly	Asp	Arg	Val	Leu	Tyr	Pro	Ala	Ser	Trp	Ile
	290					295					300				
Val	Pro	Leu	Phe	Val	Ala	Phe	Ser	Thr	Ile	Gly	Ala	Ala	Asn	Gly	Thr
305					310					315					320
Cys	Phe	Thr	Ala	Gly	Arg	Leu	Ile	Tyr	Val	Ala	Gly	Arg	Glu	Gly	His
				325					330					335	
Met	Leu	Lys	Val	Leu	Ser	Tyr	Ile	Ser	Val	Arg	Arg	Leu	Thr	Pro	Ala

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			340					345					350		
Pro	Ala	Ile	Ile	Phe	Tyr	Gly	Ile	Ile	Ala	Thr	Ile	Tyr	Ile	Ile	Pro
•		355					360			٠		365			
Gly	Asp	Ile	Asn	Ser	Leu	Val	Asn	Tyr	Phe	Ser	Phe	Ala	Ala	Trp	Leu
	370					375					380				
Phe	Tyr	Gly	Leu	Thr	Ile	Leu	Gly	Leu	Ile	Val	Met	Arg	Phe	Thr	Arg
385					390					395					400
Lys	Glu	Leu	Glu	Arg	Pro	Ile	Lys	Val	Pro	Val	Val	Ile	Pro	Val	Leu
				405					410					415	
Met	Thr	Leu	Ile	Ser	Val	Phe	Leu	Val	Leu	Ala	Pro	Ile	Ile	Ser	Lys
			420					425					430		
Pro	Thr	Trp	Glu	Tyr	Leu	Tyr	Cys	Val	Leu	Phe	Ile	Leu	Ser	Gly	Leu
		435					440			•		445	•		
Leu	Phe	Tyr	Phe	Leu	Phe	Val	His	Tyr	Lys	Phe	Gly	Trp	Ala	Gln	Lys
	450					455					460				
Ile	Ser	Lys	Pro	Ile	Thr	Met	His	Leu	Gln	Met	Leu	Met	Glu	Val	Val
465					470					475					480
Pro	Pro	Glu	Glu	Asp	Pro	Glu									
				485											
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⟨21	1> 3	75													

<212> PRT

<400> 34

<213≻ Homo sapiens

Met	Thr	Pro	Gln	Pro	Ala	Gly	Pro	Pro	Asp	Gly	Gly	Trp	Gly	Trp	Val
.1				5					10					15	
Val	Ala	Ala	Ala	Ala	Phe	Ala	Ile	Asn	Gly	Leu	Ser	Tyr	Gly	Leu	Leu
			20					25					30		
Arg	Ser	Leu	Gly	Leu	Ala	Phe	Pro	Asp	Leu	Ala	Glu	His	Phe	Asp	Arg
		35					40					45			
Ser	Ala	Gln	Asp	Thr	Ala	Trp	Ile	Ser	Ala	Leu	Ala	Leu	Ala	Val	Gln
	50					55					60				
Gln	Ala	Ala	Ser	Pro	Val	Gly	Ser	Ala	Leu	Ser	Thr	Arg	Trp	Gly	Ala
65					70					75					80
Arg	Pro	Val	Val	Met	Val	Gly	Gly	Val	Leu	Ala	Ser	Leu	Gly	Phe	Val
				85					90					95	
Phe	Ser	Ala	Phe	Ala	Ser	Gly	Leu	Leu	His	Leu	Tyr	Leu	Gly	Leu	Gly
			100					105					110		
Leu	Leu	Ala	Gly	Phe	Gly	Trp	Ala	Leu	Val	Phe	Ala	Pro	Ala	Leu	Gly
		115					120					125			
Thr	Leu	Ser	Arg	Tyr	Phe	Ser	Arg	Arg	Arg	Val	Leu	Ala	Val	Gly	Leu
	130					135					140				
Ala	Leu	Thr	Gly	Asn	Gly	Ala	Ser	Ser	Leu	Leu	Leu	Ala	Pro	Ala	Leu
145					150					155					160
Gln	Leu	Leu	Leu	Asp	Thr	Phe	Gly	Trp	Arg	Gly	Ala	Leu	Leu	Leu	Leu
				165					170					175	
Gly	Ala	Ile	Thr	Leu	His	Leu	Thr	Pro	Cys	Gly	Ala	Leu	Leu	Leu	Pro
			180					185					190		
Leu	Val	Leu	Pro	Gly	Asp	Pro	Pro	Ala	Pro	Pro	Arg	Ser	Pro	Leu	Ala

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		195					200					205			
Ala L	.eu	Gly	Leu	Ser	Leu	Phe	Thr	Arg	Arg	Ala	Phe	Ser	Ile	Phe	Ala
2	210					215				•	220				
Leu G	Gly	Thr	Ala	Leu	Val	Gly	Gly	Gly	Tyr	Phe	Val	Pro	Tyr	Val	His
225					230					235					240
Leu A	lla	Pro	Arg	Phe	Arg	Pro	Gly	Pro	Gly	Gly	Ile	Arg	Ser	Ser	Ala
				245					250					255	
Gly G	Gly	Gly	Arg	Gly	Cys	Asp	G1y	Gly	Cys	Gly	Arg	Pro	Ala	Gly	Leu
			260					265					270		
Arg V	/al	Ala	Gly	Arg	Pro	Arg	Leu	Gly	Ala	Pro	Pro	Ala	Ala	Ala	Gly
		275					280				٠	285			
Arg]	lle	Arg	Gly	Ser	Asp	Trp	Ala	Gly	Ala	Val	Gly	Gly	Gly	Ala	Gly
2	290					295					300				
Ala A	Arg	Gly	Gly	Arg	Arg	Arg	Glu	Leu	Gly	Gly	Ser	Pro	Ala	Gly	Arg
305					310					315					320
Gly (Cys	Gly	Leu	Trp	Ala	Glu	Arg	Gly	Glu	Leu	Arg	Pro	Ala	Gly	Phe
				325					330					335	
Arg (Cys	Thr	Pro	Arg	Ala	Gly	Gly	Arg	Arg	Arg	Cys	Gly	Ala	Gly	His
			340					345					350		
Arg /	Ala	Gly	Asp	Asp	Ala	Asp	Glu	Pro	Arg	Gly	Ala	Pro	Gly	Pro	Ser
		355					360					365			
Pro V	Val	Arg	Leu	Pro	Lys	Gly									
:	370					375									

⟨210⟩ 35

(211	> 35	0													
<212	> PR	T													
<213	> Ho	omo s	apie	ns		•	•		•						
<400	> 35	;													
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Pro	Glu	Trp	Gly	Gly	Phe	Glu	Glu	Asn	Ile	Gln	Gly	Gly	Gly	Ser	Ala
			20					25					30		
Val	Ile	Asp	Met	Glu	Asn	Met	Asp	Asp	Thr	Ser	Gly	Ser	Ser	Phe	Glu
		35					40					45			
Asp	Met	Gly	Glu	Leu	His	Gln	Arg	Leu	Arg	Glu	Glu	Glu	Val	Asp	Ala
	50					55					60				
Asp	Ala	Ala	Asp	Ala	Ala	Ala	Ala	Glu	Glu	Glu	Asp	Gly	Glu	Phe	Leu
65					70					75					80
Gly	Met	Lys	Gly	Phe	Lys	Gly	Gln	Leu	Ser	Arg	Gln	Val	Ala	Asp	Gln
				85					90					95	
Met	Trp	Gln	Ala	Gly	Lys	Arg	Gln	Ala	Ser	Arg	Ala	Phe	Ser	Leu	Tyr
			100					105					110		
Ala	Asn	Ile	Asp	Ile	Leu	Arg	Pro	Tyr	Phe	Asp	Val	Glu	Pro	Ala	Gln
		115					120					125			
Val	Arg	Ser	Arg	Leu	Leu	Glu	Ser	Met	Ile	Pro	Ile	Lys	Met	Val	Asn
	130					135					140				
Phe	Pro	Gln	Lys	Ile	Ala	Gly	Glu	Leu	Tyr	Gly	Pro	Leu	Met	Leu	Val
145					150					155					160
Dha	Thr	ī ou	Va 1	Δla	Tla	יים ז	Lau	Hic	C1 v	Mat	Ive	Thr	Ser	Asn	Thr

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				165					170					175	
Ile	Ile	Arg	Glu	Gly	Thr	Leu	Met	Gly	Thr	Ala	Ile	Gly	Thr	Cys	Phe
			180					185					190		
Gly	Tyr	Trp	Leu	Gly	Val	Ser	Ser	Phe	Ile	Tyr	Phe	Leu	Ala	Tyr	Leu
		195					200					205			
Cys	Asn	Ala	Gln	Ile	Thr	Met	Leu	Gln	Met	Leu	Ala	Leu	Leu	Gly	Tyr
	210					215					220				
Gly	Leu	Phe	Gly	His	Cys	Ile	Val	Leu	Phe	Ile	Thr	Tyr	Asn	Ile	His
225					230					235					240
Leu	His	Ala	Leu	Phe	Tyr	Leu	Phe	Trp	Leu	Leu	Val	Gly	Gly	Leu	Ser
				245					250					255	
Thr	Leu	Arg	Met	Val	Ala	Val	Leu	Val	Ser	Arg	Thr	Val	Gly	Pro	Thr
		•	260					265					270		
Gln	Arg	Leu	Leu	Leu	Cys	Gly	Thr	Leu	Ala	Ala	Leu	His	Met	Leu	Phe
		275					280					285			
Leu	Leu	Tyr	Leu	His	Phe	Ala	Tyr	His	Lys	Val	Val	Glu	Gly	Ile	Leu
	290					295					300				
Asp	Thr	Leu	Glu	Gly	Pro	Asn	Ile	Pro	Pro	Ile	Gln	Arg	Val	Pro	Arg
305					310					315	•				320
Asp	Ile	Pro	Ala	Met	Leu	Pro	Ala	Ala	Arg	Leu	Pro	Thr	Thr	Val	Leu
				325					330					335	
Asn	Ala	Thr	Ala	Lys	Ala	Val	Ala	Val	Thr	Leu	Gln	Ser	His		
			340					345					350		

<210> 36

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<212	2> PF	RT.													
<213	3> Ho	omo s	sapie	ens											
<400)> 36	5													
Met	Ser	Ser	Gln	Pro	Ala	Gly	Asn	Gln	Thr	Ser	Pro	Gly	Ala	Thr	Glu
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Asp	Tyr	Ser	Tyr	Gly	Ser	Trp	Tyr	Ile	Asp	Glu	Pro	Gln	Gly	Gly	Glu
			20					25					30		
Glu	Leu	G1n	Pro	Glu	Gly	Glu	Val	Pro	Ser	Cys	His	Thr	Ser	Ile	Pro
		35					40					45			
Pro	Gly	Leu	Tyr	His	Ala	Cys	Leu	Ala	Ser	Leu	Ser	Ile	Leu	Val	Leu
	50					55					60				
Leu	Leu	Leu	Ala	Met	Leu	Val	Arg	Arg	Arg	Gln	Leu	Trp	Pro	Asp	Cys
65					70					75					80
Val	Arg	Gly	Arg	Pro	Gly	Leu	Pro	Ser	Pro	Val	Asp	Phe	Leu	Ala	Gly
				85					90					95	
Asp	Arg	Pro	Arg	Ala	Val	Pro	Ala	Ala	Väl	Phe	Met	Val	Leu	Leu	Ser
			100					105					110		
Ser	Leu	Cys	Leu	Leu	Leu	Pro	Asp	Glu	Asp	Ala	Leu	Pro	Phe	Leu	Thr
		115					120					125			
Leu	Ala	Ser	Ala	Pro	Ser	Gln	Asp	Gly	Lys	Thr	Glu	Ala	Pro	Arg	Gly
	130					135					140				
Ala	Trp	Lys	Ile	Leu	Gly	Leu	Phe	Tyr	Tyr	Ala	Ala	Leu	Tyr	Tyr	Pro
145					150					155					160
Leu	Ala	Ala	Cys	Ala	Thr	Ala	Gly	His	Thr	Ala	Ala	His	Leu	Leu	Gly

				165					170					175	
Ser	Thr	Leu	Ser	Trp	Ala	His	Leu	Gly	Val	Gln	Val	Trp	Gln	Arg	Ala
			180					185					190		
Glu	Cys	Pro	G1n	Val	Pro	Lys	Ile	Tyr	Lys	Tyr	Tyr	Ser	Leu	Leu	Ala
		195					200					205			
Ser	Leu	Pro	Leu	Leu	Leu	Gly	Leu	Gly	Phe	Leu	Ser	Leu	Trp	Tyr	Pro
	210					215					220				
Val	Gln	Leu	Val	Arg	Ser	Phe	Ser	Arg	Arg	Thr	Gly	Ala	Gly	Ser	Lys
225					230					235					240
Gly	Leu	Gln	Ser	Ser	Tyr	Ser	Glu	Glu	Tyr	Leu	Arg	Asn	Leu	Leu	Cys
				245					250					255	
Arg	Lys	Lys	Leu	Gly	Ser	Ser	Tyr	His	Thr	Ser	Lys	His	Gly	Phe	Leu
			260				•	265					270		
Ser	Trp	Ala	Arg	Val	Cys	Leu	Arg	His	Cys	Ile	Tyr	Thr	Pro	Gln	Pro
		275					280					285			
Gly	Phe	His	Leu	Pro	Leu	Lys	Leu	Val	Leu	Ser	Ala	Thr	Leu	Thr	Gly
	290					295					300				
Thr	Ala	Ile	Tyr	Gln	Val	Ala	Leu	Leu	Leu	Leu	Val	Gly	Val	Val	Pro
305			. •		310					315					320
Thr	Ile	Gln	Lys	Val	Arg	Ala	Gly	Val	Thr	Thr	Asp	Val	Ser	Tyr	Leu
				325					330					335	
Leu	Ala	Gly	Phe	G1y	Ile	Val	Leu	Ser	Glu	Asp	Lys	G1n	Glu	Val	Val
			340)				345					350		
Glu	Leu	Val	Lys	His	His	Leu	Trp	Ala	Leu	Glu	Val	Cys	Tyr	Ile	Ser
		355					360					365	;		

Ala	Leu	Val	Leu	Ser	Cys	Leu	Leu	Thr	Phe	Leu	Val	Leu	Met	Arg	Ser
	370·					375					380		•		
Leu	Val	Thr	His	Arg	Thr	Asn	Leu	Arg	Ala	Leu	His	Arg	Gly	Ala	Ala
385					390					395					400
Leu	Asp	Leu	Ser	Pro	Leu	His	Arg	Ser	Pro	His	Pro	Ser	Arg	Gln	Ala
				405					410					415	
Ile	Phe	Cys	Trp	Met	Ser	Phe	Ser	Ala	Tyr	Gln	Thr	Ala	Phe	Ile	Cys
			420					425					430		
Leu	Gly	Leu	Leu	Val	Gln	Gln	Ile	Ile	Phe	Phe	Leu	Gly	Thr	Thr	Ala
		435					440					445			
Leu	Ala	Phe	Leu	Val	Leu	Met	Pro	Val	Leu	His	Gly	Arg	Asn	Leu	Leu
	450					455					460				
Leu	Phe	Arg	Ser	Leu	Glu	Ser	Ser	Trp	Pro	Phe	Trp	Leu	Thr	Leu	Ala
465					470					475					480
Leu	Ala	Val	Ile	Leu	Gln	Asn	Met	Ala	Ala	His	Trp	Val	Phe	Leu	Glu
				485					490					495	
Thr	His	Asp	Gly	His	Pro	Gln	Leu	Thr	Asn	Arg	Arg	Val	Leu	Tyr	Ala
			500	•				505					510		
Ala	Thr	Phe	Leu	Leu	Phe	Pro	Leu	Asn	Val	Leu	Val	Gly	Ala	Met	Val
		515	,				520	ı				525	;		
Ala	Thr	Trp	Arg	Yal	Leu	Leu	Ser	Ala	Leu	Tyr	Asn	Ala	Ile	His	Leu
	530)				535	;				540)			
Gly	Gln	Met	. Asp	Leu	Ser	Leu	Leu	Pro	Pro	Arg	Ala	Ala	Thr	Leu	Asp
545	i				550)				555	,				560
Pro	Gly	Туз	Tyı	Thr	Tyr	Arg	, Asn	Phe	Leu	Lys	Ile	Glu	ı Val	Ser	·Gln

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Ser His Pro Ala Met Thr Ala Phe Cys Ser Leu Leu Leu Gln Ala Gln 580 . Ser Leu Leu Pro Arg Thr Met Ala Ala Pro Gln Asp Ser Leu Arg Pro Gly Glu Glu Asp Glu Gly Met Gln Leu Leu Gln Thr Lys Asp Ser Met Ala Lys Gly Ala Arg Pro Gly Ala Ser Arg Gly Arg Ala Arg Trp Gly Leu Ala Tyr Thr Leu Leu His Asn Pro Thr Leu Gln Val Phe Arg Lys Thr Ala Leu Leu Gly Ala Asn Gly Ala Gln Pro <210> 37 <211> 464 <212> PRT <213> Homo sapiens <400> 37 Met Ile Val Cys Leu Leu Phe Met Met Ile Leu Leu Ala Lys Glu Val Gln Leu Val Asp Gln Thr Asp Ser Pro Leu Leu Ser Leu Leu Gly Gln Thr Ser Ser Leu Ser Trp His Leu Val Asp Ile Val Ser Tyr Gln Ser

Val	Leu	Ser	Tyr	Phe	Ser	Ser	His	Tyr	Pro	Pro	Ser	Ile	Ile	Leu	Ala
	50					55					60				
Lys	Glu	Ser	Tyr	Ala	Glu	Leu	Ile	Met	Lys	Leu	Leu	Lys	Val	Ser	Ala
65					70					75					80
Gly	Leu	Ser	Ile	Pro	Thr	Asp	Ser	Gln	Lys	His	Leu	Asp	Ala	Val	Pro
				85					90					95	
Lys	Cys	Gln	Ala	Phe	Thr	His	Gln	Met	Val	Gln	Phe	Leu	Ser	Thr	Leu
			100					105					110		
Glu	Gln	Asn	Gly	Lys	Ile	Thr	Leu	Ala	Val	Leu	Glu	Gln	Glu	Met	Ser
		115					120					125			
Lys	Leu	Leu	Asp	Asp	Ile	Ile	Val	Phe	Asn	Pro	Pro	Asp	Met	Asp	Ser
	130)				135					140			٠	
Glr	Thi	Arg	His	Met	. Ala	Leu	Ser	Ser	Leu	Phe	Met	Glu	Val	Leu	Met
145	5		•		150)				155					160
Met	t Me	t Asr	. Asr	n Ala	a Thr	· Ile	Pro	Thr	Ala	Glu	Phe	Leu	Arg	Gly	Ser
				168	5				170)				175	
Ile	e Ar	g Thi	Tr	o Ile	e Gly	Glr	ı Lys	Met	His	Gly	Leu	Val	. Val	Leu	Pro
			180)				185	j				190)	
Le	u Le	u Thi	r Ala	a Ala	a Cys	s Glr	n Ser	Leu	ı Ala	a Ser	· Val	Are	g His	s Met	Ala
		19	5				200)				205	5		
Gl	u Th	r Th	r Gl	u Al	a Cy:	s Il	e Thi	r Ala	з Туз	r Phe	e Lys	s Glu	ı Sei	r Pro	Leu
	21	0				21	5				220)			
As	n Gl	n As	n Se	r Gl	y Tr	p Gl	y Pr	o Ile	e Le	u Va	1 Se	r Le	u Gli	n Va	l Pro
22	:5				23	0				23	5				240
Gl	u Le	eu Th	r Me	t Gl	u Gl	u Ph	e Le	u G1:	n Gl	u Cy	s Le	u Th	r Le	u Gl	y Ser

				245					250					255	
Tyr	Leu	Thr	Leu	Tyr	Val	Tyr	Leu	Leu	Gln	Cys	Leu	Asn	Ser	Glu	Gln
			260			•		265					270		
Thr	Leu	Arg	Asn	Glu	Met	Lys	Val	Leu	Leu	Ile	Leu	Ser	Lys	Trp	Leu
		275					280					285			
Glu	Gln	Val	Tyr	Pro	Ser	Ser	Val	Glu	Glu	Glu	Ala	Lys	Leu	Phe	Leu
	290					295		-			300				
Trp	Trp	His	G1n	Val	Leu	Gln	Leu	Ser	Leu	Ile	Gln	Thr	Glu	Gln	Asn
305	•				310					315					320
Asp	Ser	Val	Leu	Thr	Glu	Ser	Val	Ile	Arg	Ile	Leu	Leu	Leu	Val	Gln
				325					330					335	
Ser	Arg	Gln	Asn	Leu	Val	Ala	Glu	Glu	Arg	Leu	Ser	Ser	Gly	Ile	Leu
			340					345					350		
Gly	Ala	Ile	Gly	Phe	Gly	Arg	Lys	Ser	Pro	Leu	Ser	Asn	Arg	Phe	Arg
		355					360					365			
Val	Val	Ala	Arg	Ser	Met	Ala	Ala	Phe	Leu	Ser	Val	Gln	Val	Pro	Met
	370					375					380				
Glu	Asp	Gln	Ile	Arg	Leu	Arg	Pro	Gly	Ser	Glu	Leu	His	Leu	Thr	Pro
385					390		•			395					400
Lys	Ala	Gln	Gln	Ala	Leu	Asn	Ala	Leu	Glu	Ser	Met	Ala	Ser	Ser	Lys
				405					410					415	
Gln	Tyr	Val	Glu	Tyr	Gln	Asp	Gln	Ile	Leu	Gln	Ala	Thr	Gln	Phe	Ile
			420					425					430		
Arg	His	Pro	Gly	His	Cys	Leu	Gln	Asp	Gly	Lys	Ser	Phe	Leu	Ala	Leu
		435					440					445			

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Leu	Val	Asn	Cys	Leu	Tyr	Pro	Glu	Val	His	Tyr	Leu	Asp	His	Ile	Arg
	450					455					460				
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<400	> 38	3													
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Leu	Leu	Gly	Thr	Ala	Ala	Gly	Leu	Gly	Phe	Leu	Cys	Leu	Leu	Tyr	Ser
			. 20					25					30		
Gln	Arg	Trp	Lys	Arg	Thr	Gln	Ärg	His	Gly	Arg	Ser	Gln	Ser	Leu	Pro
		35					40					45			
Asn	Ser	Leu	Asp	Tyr	Thr	Gln	Thr	Ser	Asp	Pro	Gly	Arg	His	Val	Met
	50					55					60				
Leu	Leu	Arg	Ala	Val	Pro	Gly	Gly	Ala	Gly	Asp	Ala	Ser	Val	Leu	Pro
65					70					75					80
Ser	Leu	Pro	Arg	Glu	Gly	Gln	Glu	Lys	Val	Leu	Asp	Arg	Leu	Asp	Phe
				85					90					95	
Val	Leu	Thr	Ser	Leu	Val	Ala	Leu	Arg	Arg	Glu	Val	Glu	Glu	Leu	Arg
			100					105					110		
Ser	Ser	Leu	Arg	Gly	Leu	Ala	Gly	Glu	Ile	Val	Gly	Glu	Val	Arg	Cys
		115					120					125			

 $\hbox{His Met Glu Glu Asn Gln Arg Val Ala Arg Arg Arg Phe Pro Phe } \\$

	130					135					140				
Val	Arg	Glu	Arg	Ser	Asp	Ser	Thr	Gly	Ser	Ser	Ser	Val	Tyr	Phe	Thr
145	•				150					155					160
Ala	Ser	Ser	Gly	Ala	Thr	Phe	Thr	Asp	Ala	Glu	Ser	Glu	Gly	Gly	Tyr
				165					170					175	
Thr	Thr	Ala	Asn	Ala	Glu	Ser	Asp	Asn	Glu	Arg	Asp	Ser	Asp	Lys	Glu
		•	180					185					190		
Ser	Glu	Asp	Gly	Glu	Asp	Glu	Val	Ser	Cys	Glu	Thr	Val	Lys	Met	Gly
		195					200					205			
Arg	Lys	Asp	Ser	Leu	Asp	Leu	Glu	Glu	Glu	Ala	Ala	Ser	Gly	Ala	Ser
	210					215					220				
Ser	Ala	Leu	Glu	Ala	Gly	Gly	Ser	Ser	Gly	Leu	Glu	Asp	Val	Leu	Pro
225					230					235					240
Leu	Leu	Gln	Gln	Ala	Asp	Glu	Leu	His	Arg	Gly	Asp	Glu	Gln	Gly	Lys
				245					250					255	
Arg	Glu	Gly	Phe	Gln	Leu	Leu	Leu	Asn	Asn	Lys	Leu	Val	Tyr	Gly	Ser
			260					265					270		
Arg	Gln	Asp	Phe	Leu	Trp	Arg	Leu	Ala	Arg	Ala	Tyr	Ser	Asp	Met	Cys
		275	•				280					285	j		
Glu	Leu	Thr	Glu	Glu	Val	Ser	Glu	Lys	Lys	Ser	Tyr	Ala	Leu	Asp	Gly
	290)				295	;				300)			
Lys	Glu	Glu	Ala	Glu	ı Ala	Ala	Leu	Glu	Lys	Gly	Asp	Glu	Ser	Ala	Asp
305	;				310)				315	5				320
Cys	His	Leu	Trp	Туг	. Ala	a Val	Leu	Cys	Gly	Glr	ı Lev	ı Ala	a Glu	ı His	Glu
				329	5				330)				335	5

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Ser Ile Gln Arg Arg Ile Gln Ser Gly Phe Ser Phe Lys Glu His Val Asp Lys Ala Ile Ala Leu Gln Pro Glu Asn Pro Met Ala His Phe Leu Leu Gly Arg Trp Cys Tyr Gln Val Ser His Leu Ser Trp Leu Glu Lys Lys Thr Ala Thr Ala Leu Leu Glu Ser Pro Leu Ser Ala Thr Val Glu Asp Ala Leu Gln Ser Phe Leu Lys Ala Glu Glu Leu Gln Pro Gly Phe Ser Lys Ala Gly Arg Val Tyr Ile Ser Lys Cys Tyr Arg Glu Leu Gly Lys Asn Ser Glu Ala Arg Trp Trp Met Lys Leu Ala Leu Glu Leu Pro Asp Val Thr Lys Glu Asp Leu Ala Ile Gln Lys Asp Leu Glu Glu Leu Glu Val Ile Leu Arg Asp

<210> 39

<211> 243

<212> PRT

<213> Homo sapiens

<400> 39

Met Glu Gln Gly Ser Gly Arg Leu Glu Asp Phe Pro Val Asn Val Phe

1				5					10					15	
Ser	Val	Thr	Pro	Tyr	Thr	Pro	Ser	Thr	Ala	Asp	Ile	Gln	Val	Ser	Asp
			20					25					30	•	
Asp	Asp	Lys	Ala	Gly	Ala	Thr	Leu	Leu	Phe	Ser	Gly	Ile	Phe	Leu	Gly
		35					40					45			
Leu	Val	Gly	Ile	Thr	Phe	Thr	Val	Met	Gly	Trp	Ile	Lys	Tyr	Gln	G1 y
	50					55					60				
Val	Ser	His	Phe	Glu	Trp	Thr	Gln	Leu	Leu	Gly	Pro	Val	Leu	Leu	Ser
65					70					75					80
Val	Gly	Val	Thr	Phe	Ile	Leu	Ile	Ala	Val	Cys	Lys	Phe	Lys	Met	Leu
				85					90					95	
Ser	Cys	Gln	Leu	Cys	Lys	Glu	Ser	Glu	Glu	Arg	Val	Pro	Asp	Ser	Glu
			100					105			•		110		
Gln	Thr	Pro	Gly	Gly	Pro	Ser	Phe	Val	Phe	Thr	Gly	Ile	Asn	Gln	Pro
		115					120					125			
Ile	Thr	Phe	His	Gly	Ala	Thr	Val	Val	Gln	Tyr	Ile	Pro	Pro	Pro	Tyr
	130					135					140				
Gly	Ser	Pro	Glu	Pro	Met	Gly	Ile	Asn	Thr	Ser	Tyr	Leu	Gln	Ser	Val
145					150					155					160
Val	Ser	Pro	Cys	Gly	Leu	Ile	Thr	Ser	Gly	Gly	Ala	Ala	Ala	Ala	Met
				165					170					175	
Ser	Ser	Pro	Pro	Gln	Tyr	Tyr	Thr	Ile	Tyr	Pro	Gln	Asp	Asn	Ser	Ala
			180					185					190		
Phe	Val	Val	Asp	Glu	Gly	Cys	Leu	Ser	Phe	Thr	Asp	Gly	Gly	Asn	His
		195					200					205			

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Arg Pro Asn Pro Asp Val Asp Gln Leu Glu Glu Thr Gln Leu Glu Glu Glu Ala Cys Ala Cys Phe Ser Pro Pro Pro Tyr Glu Glu Ile Tyr Ser Leu Pro Arg <210> 40 <211> 270 <212> PRT <213> Homo sapiens <400> 40 Met Ala Gly Ala Glu Asp Trp Pro Gly Gln Gln Leu Glu Leu Asp Glu Asp Glu Ala Ser Cys Cys Arg Trp Gly Ala Gln His Ala Gly Ala Arg Glu Leu Ala Ala Leu Tyr Ser Pro Gly Lys Arg Leu Gln Glu Trp Cys Ser Val Ile Leu Cys Phe Ser Leu Ile Ala His Asn Leu Val His Leu Leu Leu Leu Ala Arg Trp Glu Asp Thr Pro Leu Val Ile Leu Gly Val Val Ala Gly Ala Leu Ile Ala Asp Phe Leu Ser Gly Leu Val His Trp Gly Ala Asp Thr Trp Gly Ser Val Glu Leu Pro Ile Val Gly Lys Ala

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Phe	Ile	Arg	Pro	Phe	Arg	Glu	His	His	Ile	Asp	Pro	Thr	Ala	Ile	Thr
		115					120					125			
Arg	His	Asp	Phe	Ile	Glu	Thr	Asn	Gly	Asp	Asn	Cys	Leu	Val	Thr	Leu
	130					135					140				
Leu	Pro	Leu	Leu	Asn	Met	Ala	Tyr	Lys	Phe	Arg	Thr	His	Ser	Pro	Glu
145					150					155					160
Ala	Leu	Glu	Gln	Leu	Tyr	Pro	Trp	Glu	Cys	Phe	Val	Phe	Cys	Leu	Ile
				165					170					175	
Ile	Phe	Gly	Thr	Phe	Thr	Asn	G1n	Ile	His	Lys	Trp	Ser	His	Thr	Tyr
			180					185					190		
Phe	Gly	Leu	Pro	Arg	Trp	'Val	Thr	Leu	Leu	Gln	Asp	Trp	His	Val	Ile
٠		195					200					205		•	
Leu	Pro	Arg	Lys	His	His	Arg	Ile	His	His	Val	Ser	Pro	His	Glu	Thr
	210					215					220				
Tyr	Phe	Cys	Ile	Thr	Thr	Gly	Trp	Leu	Asn	Tyr	Pro	Leu	Glu	Lys	Ile
225					230					235					240
Gly	Phe	Trp	Arg	Arg	Leu	Glu	Asp	Leu	Ile	Gln	Gly	Leu	Thr	Gly	Glu
				245					250					255	
Lys	Pro	Arg	Ala	Asp	Asp	Met	Lys	Trp	Ala	Gln	Lys	Ile	Lys		
			260					265					270		

<210> 41

<211> 1131

<212> DNA

<213> Homo sapiens

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⟨400⟩ 41

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180	cgcccgcggc	ccgtacgctg	gccctgcgct	cttcttcggc	tgctgcccat	ctcatggcgc
240	cttccccatc	atgccgcccg	accagccggg	tgaaacaatc	cagacatgcc	aagaatgctt
300	ccaggagtac	aaatattctc	ctcttttca	ggggctctac	gcacactctt	atcgccagct
360	gtcccacacc	tcctggccct	gtgctgggaa	gtatttcttc	tgctgtccat	atcaacctcc
420	gtaccagctg	caaatcgaca	gccagctttc	gtttttcca	tcatgaataa	atcagcccct
480	atttgacacc	tcaattatga	gaagagatca	ggaaaacaag	agggttctgg	ctcttcacac
540	gctgaggaag	tctggtacct	atcgttggcg	cctgagcagc	tgtgcctggg	aaggacctgg
600	agagctcctg	ttaatggagt	gccttctccc	ttttggcctg	ccaacaacct	cactggattg
660	ctacgatgtc	gactcttcat	ctgctgggcg	tggctgcatc	atgtcagcac	cacctcaaca
720	ggcaccaata	agtccttcga	acagtggcca	tgtgatggtg	ttggcaccaa	ttctgggtat
780	ctttgccatg	aagcaaacaa	aaaggcctcg	tctgctggag	ttccccagga	aaattggtgt
840	gcgctttgac	ccttgctgct	atcttcattg	cattccaggg	gagatgtcgt	ctgggacttg
900	ctacatcttc	gctttgcagc	ttctacacca	ccacacctac	agaagaatac	atcagcttga
960	tgccctccta	atgctcagcc	atcttcaagc	catcatgcac	ttaccatctt	ggcctgggcc
1020	gggagaagtg	cgctggccaa	gtcctggtgg	cggttttcct	ccgcctgcat	tacctggtcc
1080	gacagaatcc	cagcggcagt	cctaaggatc	ggagtcaaat	tcagttatga	acagagatgt
1131	а	agaaagagaa	gggctggaga	agcatcgaag	cagaggcatc	aaagagggaa

<210> 42

⟨211⟩ 243

<212> DNA

<213> Homo sapiens

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<	4	U	Λ	>	42
`	7	v	v	/	74

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<210> 43

(211) 1461

<212> DNA

<213> Homo sapiens

<400> 43

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ccc	ctggtga	cggcgtgcta	catcctcatg	aacgtgtcct	acttcaccgt	gatgactgcc	840
300	gaactcc	tgcagtccca	ggcggtggct	gtgacatttg	gtgaccgtgt	tctctatcct	. 900
gc1	ttcttgga	tcgttccact	ttttgtggca	ttttcaacca	tcggtgctgc	taacgggacc	960
tgo	cttcacag	cgggcagact	catttacgtg	gcgggccggg	agggtcacat	gctcaaagtg	1020
eti	ttcttaca	tcagcgtcag	gcgcctcact	ccagcccccg	ccatcatctt	ttatggtatc	1080
ata	agcaacga	tttatatcat	ccctggtgac	ataaactcgt	tagtcaatta	tttcagcttt	1140
gco	egcatggc	tgttttatgg	cctgacgatt	ctaggactca	tcgtgatgag	atttacaagg	1200
aaa	agagctgg	aaaggcctat	caaggtgccc	gtagtcattc	ccgtcttgat	gacactcatc	1260
tci	tgtgtttt	tggttctggc	tccaatcatc	agcaagccca	cctgggagta	cctctactgt	1320
gtį	gctgttta	tattaagcgg	ccttttattt	tacttcctgt	ttgtccacta	caagtttgga	1380
tg	ggctcaga	aaatctcaaa	gccgattacc	atgcaccttc	agatgctaat	ggaagtggtc	1440
cca	accggagg	aagaccctga	g				1461

⟨210⟩ 44

⟨211⟩ 1125

<212> DNA

<213≻ Homo sapiens

⟨400⟩ 44

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1	gccttcgcga	taaacgggct	gtcctacggg	ctgctgcgct	cgctgggcct	tgccttccct	120
1	gaccttgccg	agcactttga	ccgaagcgcc	caggacactg	cgtggatcag	cgccctggcc	180
(ctggccgtgc	agcaggcagc	cagccccgtg	ggcagcgccc	tgagcacgcg	ctggggggcc	240
(cgccccgtgg	tgatggttgg	gggcgtcctc	gcctcgctgg	gcttcgtctt	ctcggctttc	300
1	gccagcggtc	tgctgcatct	ctacctcggc	ctgggcctcc	tcgctggctt	tggttgggcc	360
,	ctggtgttcg	ccccgccct	aggcaccctc	tcgcgttact	tctcccgccg	tcgagtcttg	420

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gcg	gtggggc	tggcgctcac	cggcaacggg	gcctcctcgc	tgctcctggc	gcccgccttg	480
cag	cttctcc	tcgatacttt	cggctggcgg	ggcgctctgc	tcctcctcgg	cgcgatcacc	540
cto	cacctca	cccctgtgg	cgccctgctg	ctacccctgg	tccttcctgg	agaccccca	600
gcc	ccaccgc	gtagtcccct	agctgccctc	ggcctgagtc	tgttcacacg	ccgggccttc	660
tca	atctttg	ctctaggcac	agccctggtt	gggggcgggt	acttcgttcc	ttacgtgcac	720
tte	gctcccc	gctttagacc	ggggcctggg	gggatacgga	gcagcgctgg	tggtggccgt	780
ggo	tgcgatg	ggggatgcgg	gcgcccggct	ggtctgcggg	tggctggcag	accaaggctg	840
ggt	gcccctc	ccgcggctgc	tggccgtatt	cggggctctg	actgggctgg	ggctgtgggt	900
ggt	ggggctg	gtgcccgtgg	tgggcggcga	agagagctgg	gggggtcccc	tgctggccgc	960
ggo	tgtggcc	tatgggctga	gcgcggggag	ttacgccccg	ctggttttcg	gtgtactccc	1020
cgg	gctggtg	ggcgtcggag	gtgtggtgca	ggccacaggg	ctggtgatga	tgctgatgag	1080
cct	cgggggg	ctcctgggcc	ctcccctgtc	aggcttccta	aggga .		1125

⟨210⟩ 45

<211> 1050

<212> DNA

<213> Homo sapiens

<400> 45

93 /307

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ggcaccctg	a tgggcacagc	cattggcacc	tgcttcggct	actggctggg	agtctcatcc	600
ttcatttac	t tccttgccta	cctgtgcaac	gcccagatca	ccatgctgca	gatgttggca	660
ctgctgggc	t atggcctctt	tgggcattgc	attgtcctgt	tcatcaccta	taatatccac	720
ctccacgcc	c tettetacet	cttctggctg	ttggtgggtg	gactgtccac	actgcgcatg	780
gtagcagtg	t tggtgtctcg	gaccgtgggc	cccacacagc	ggctgctcct	ctgtggcacc	840
ctggctgcc	c tacacatgct	cttcctgctc	tatctgcatt	ttgcctacca	caaagtggta	900
gaggggatc	c tggacacact	ggagggcccc	aacatcccgc	ccatccagag	ggtccccaga	960
gacatccct	g ccatgctccc	tgctgctcgg	cttcccacca	ccgtcctcaa	cgccacagcc	1020
aaagctgtt	g cggtgaccct	gcagtcacac				1050

<210> 46

<211> 2001

<212> DNA

<213> Homo sapiens

<400> 46

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ctttggtacc	ctgtgcagct	ggtgagaagc	ttcagccgta	ggacaggagc	aggctccaag	720
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ggaagcagct	accacacctc	caagcatggc	ttcctgtcct	gggcccgcgt	ctgcttgaga	840
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acactgacag	ggacggccat	ttaccaggtg	gccctgctgc	tgctggtggg	cgtggtaccc	960
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gctctggaag	tgtgctacat	ctcagccttg	gtcttgtcct	gcttactcac	cttcctggtc	1140
ctgatgcgct	cactggtgac	acacaggacc	aaccttcgag	ctctgcaccg	aggagctgcc	1200
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gccatccaco	ttggccagat	ggacctcagc	ctgctgccac	cgagagccgc	cactctcgac	1680
cccggctact	t acacgtaccg	aaacttcttg	, aagattgaag	tcagccagtc	gcatccagcc	1740
atgacagcc	t tetgeteect	gctcctgcaa	gcgcagagco	: tcctacccag	gaccatggca	1800
gcccccag	g acagootoag	accaggggag	gaagacgaag	g ggatgcagct	gctacagaca	1860
aaggactcc	a tggccaaggg	agctaggcc	ggggccagco	gcggcaggg	ctcgctggggt	1920
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ggtgccaatg gtgcccagcc c 2001

<210> 47

⟨211⟩ 1392

<212> DNA

<213> Homo sapiens

<400> 47

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tcgcctttgt	ctaacaggtt	ccgagtggtt	gcccgaagca	tggctgcctt	cctttcagtt	1140
caggttccta	tggaagatca	gatccgtttg	aggcctggct	ctgaattaca	tctgaccccc	1200
aaagctcagc	aggctctgaa	tgctcttgaa	tccatggcat	caagtaagca	gtatgttgaa	1260
taccaggatc	aaatattgca	agccacccaa	tttataaggc	atcctggcca	ttgccttcaa	1320
gatgggaaaa	gcttcttggc	tcttctcgtt	aactgtctgt	atccagaagt	gcattatttg	1380
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⟨210⟩ 48

<211> 1410

<212> DNA

<213> Homo sapiens

<400> 48

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cagctgctgc	tcaacaacaa	gctggtgtat	ggaagccggc	aggactttct	ctggcgcctg	840
gcccgagcct	acagtgacat	gtgtgagctc	actgaggagg	tgagcgagaa	gaagtcatat	900
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gaaaacccca	tggctcactt	tcttcttggc	aggtggtgct	atcaggtctc	tcacctgagc	1140
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<210> 49

<211> 729

<212> DNA

<213> Homo sapiens

<400> 49

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gtgagcccct	gcggcctcat	aacctctgga	ggggcagcag	ccgccatgtc	aagtcctcct	540
caatactaca	ccatctaccc	tcaagataac	tctgcatttg	tggttgatga	gggctgcctt	600
tctttcacgg	acggtggaaa	tcacaggccc	aatcctgatg	ttgaccagct	agaagagaca	660
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<210> 50

⟨211⟩ 810

<212> DNA

<213> Homo sapiens

<400> 50

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		Met Asp Ser	Ala Leu Ser
		1	5
gat ccg cat	aac ggc agt gcc gag	gca ggc ggc ccc acc	aac agc act 163
Asp Pro His	Asn Gly Ser Ala Glu	Ala Gly Gly Pro Thr	Asn Ser Thr
	10	15	20
acg cgg ccg	cct tcc acg ccc gag	ggc atc gcg ctg gcc	tac ggc agc 211
Thr Arg Pro	Pro Ser Thr Pro Glu	Gly Ile Ala Leu Ala	Tyr Gly Ser
25	30	. 35	
ctc ctg ctc	atg gcg ctg ctg ccc	e atc ttc ttc ggc gcc	ctg cgc tcc 259
Leu Leu Leu	Met Ala Leu Leu Pro	o Ile Phe Phe Gly Ala	Leu Arg Ser
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gta cgc tgc	gcc cgc ggc aag aa	t gct tca gac atg cct	gaa aca atc 307
Val Arg Cys	Ala Arg Gly Lys As	n Ala Ser Asp Met Pro	Glu Thr Ile
55	60	65	70 _,

acc	agc	cgg	gat	gcc	gcc	cgc	ttc	ссс	atc	atc	gcc	agc	tgc	aca	ctc	355
Thr	Ser	Arg	Asp	Ala	Ala	Arg	Phe	Pro	Ile	Ile	Ala	Ser	Cys	Thr	Leu	
				75					80				•	85		
ttg	ggg	ctc	tac	ctc	ttt	ttc	aaa	ata	ttc	tcc	cag	gag	tac	atc	aac	403
Leu	Gly	Leu	Tyr	Leu	Phe	Phe	Lys	Ile	Phe	Ser	Gln	Glu	Tyr	Ile	Asn	
			90					95					100		•	
ctc	ctg	ctg	tcc	atg	tat	ttc	ttc	gtg	ctg	gga	atc	ctg	gcc	ctg	tcc	451
Leu	Leu	Leu	Ser	Met	Tyr	Phe	Phe	Val	Leu	Gly	Ile	Leu	Ala	Leu	Ser	
		105					110					115				
cac	acc	atc	agc	ссс	ttc	atg	aat	aag	ttt	ttt	cca	gcc	agc	ttt	cca	499
His	Thr	Ile	Ser	Pro	Phe	Met	Asn	Lys	Phe	Phe	Pro	Ala	Ser	Phe	Pro	
	120					125					130					
aat	cga	cag	tac	cag	ctg	ctc	ttc	aca	cag	ggt	tct	ggg	gaa	aac	aag	547
Asn	Arg	Gln	Tyr	Gln	Leu	Leu	Phe	Thr	Gln	Gly	Ser	Gly	Glu	Asn	Lys	
135					140	•				145					150	
gaa	gag	ato	atc	aat	tat	gaa	ttt	gac	acc	aag	gac	ctg	gtg	tgo	ctg	595
Glu	Glu	Ile	Ile	Asn	Tyr	Glu	Phe	Asp	Thr	Lys	Asp	Leu	Va]	Cys	Leu	
				155	•				160)				165	5	
ggc	ctg	gʻago	ago	ato	gtt	ggc	gto	tgg	tac	ctg	ctg	agg	g aag	z cac	tgg	643
Gly	Leu	ı Ser	Ser	Ile	· Val	Gly	v Val	l Trp	Туг	Leu	ı Leu	Arg	χ Ly:	s His	s Trp	
			170)				175	5				180	0		
att	gco	e aac	aac	ctt	tt1	t ggo	cti	g gc	c tto	tco	c cti	aa1	t gg:	a gt	a gag	691
Ile	Ala	a Ası	n Asr	Leu	ı Phe	e Gly	/ Le	u Ala	a Phe	e Sei	r Lei	ı Ası	n Gl	y Va	l Glu	
		18	5				19	0				19	5			
			o ct/		- 99	t ot	- aa	C SC	t oo	c ta	c at	e et	g ct	g gg	c gga	739

Leu	Leu	His	Leu	Asn	Asn	Val	Ser	Thr	Gly	Cys	Ile	Leu	Leu	Gly	Gly	
	200					205					210					
ctc	ttc	atc	tac	gat	gtc	ttc	tgg	gta	ttt	ggc	acc	aat	gtg	atg	gtg	787
Leu	Phe	Ile	Tyr	Asp	Val	Phe	Trp	Val	Phe	Gly	Thr	Asn	Val	Met	Val	
215					220					225					230	
aca	gtg	gcc	aag	tcc	ttc	gag	gca	cca	ata	aaa	ttg	gtg	ttt	ccc	cag	835
Thr	Val	Ala	Lys	Ser	Phe	Glu	Ala	Pro	Ile	Lys	Leu	Val	Phe	Pro	Gln	
				235					240					245		
gat	ctg	ctg	gag	aaa	ggc	ctc	gaa	gca	aac	aac	ttt	gcc	atg	ctg	gga	883
Asp	Leu	Leu	Glu	Lys	Gly	Leu	Glu	Ala	Asn	Asn	Phe	Ala	Met	Leu	Gly	
			250					255					260			
ctt	gga	gat	gtc	gtc	atţ	cca	ggg	atc	ttc	att	gcc	ttg	ctg	ctg	cgc	931
Leu	Gly	Asp	Val	Val	Ile	Pro	Gly	Ile	Phe	Ile	Ala	Leu	Leu	Leu	Arg	
		265					270					275				
ttt	gac	atc	agc	ttg	aag	aag	aat	acc	cac	acc	tac	ttc	tac	acc	agc	979
Phe	. Asp	Ile	Ser	Leu	Lys	Lys	Asn	Thr	His	Thr	Tyr	Phe	Туг	Thr	Ser	
	280)				285					290)				
ttt	gca	g gcc	tac	ato	ttc	ggc	ctg	ggc	ctt	acc	ato	ttc	ato	ate	g cac	1027
Phe	e Ala	a Ala	Tyr	Ile	Phe	Gly	Leu	Gly	Leu	Thr	Ile	Phe	: Ile	e Met	His	
295	5				300	+				305	5				310	
ato	t tt	c aag	g cat	gct	; cag	cct	gcc	cto	cta	tac	cts	ggto	cc	c gc	tgc	1075
Ile	e Ph	e Lys	s His	s Ala	Gln	Pro	Ala	Leu	ı Leu	Туз	Leu	ı Val	Pre	o Ala	a Cys	
				315	5				320)				32	5	
at	c gg	t tt	t cc1	t gto	ctg	gtg	g gcg	g ctg	g gcc	aag	g gg	a gaa	a gt	g ac	a gag	1123
11	e Gl	y Pho	e Pro	o Val	l Lei	ı Val	l Ala	a Let	ı Ala	a Ly:	s Gl	y Gl	u Va	1 Th	r Glu	

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			220					335					340			
			330					333					340			
atg	ttc	agt	tat	gag	gag	tca	aat	cct	aag	gat	cca	gcg	gca	gtg	aca	1171
Met	Phe	Ser	Tyr	Glu	Glu	Ser	Asn	Pro	Lys	Asp	Pro	Ala	Ala	Val	Thr	
		345					350					355				
gaa	tcc	aaa	gag	gga	aca	gag	gca	tca	gca	tcg	aag	ggg	ctg	gag	aag	1219
Glu	Ser	Lys	Glu	Gly	Thr	Glu	Ala	Ser	Ala	Ser	Lys	Gly	Leu	Glu	Lys	
	360					365					370					
aaa	gag	aaa	tg a	atgc	agct	gg t	gccc	gago	c tc	tcag	ggcc	aga	ccag	aca		1270
Lys	Glu	Lys											•			
375																
gat	gggg	gct	gggc	ccac	ac a	ggcg	tgca	с сд	gtag	aggg	cac	agga	ggc	caag	ggcagc	1330
tcc	agga	cag	ggca	gggg	gc a	gcag	gatạ	c ct	ccag	ccag	gcc	tctg	tgg	cctc	tgtttc	1390
ctt	ctcc	ctt	tctt	ggcc	ct c	ctct	gctc	c to	ссса	cacc	ctg	cagg	caa	aaga	aacccc	1450
cag	ctto	ccc	cctc	cccg	gg a	igcca	ggtg	g ga	aaag	tggg	tgt	gatt	ttt	agat	tttgta	1510
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<210> 52

<211> 1713

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> (89)...(334)

<400> 52

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							, N	let 1	Thr A	la F	lis S	Ser F	he A	Ala I	_eu		
			٠					1				5					
ccg	gtc	atc	atc	ttc	acc	acg	ttc	tgg	ggc	ctc	gtc	ggc	atc	gcc	ggg		160
Pro	Val	Ile	·Ile	Phe	Thr	Thr	Phe	Trp	Gly	Leu	Val	Gly	Ile	Ala	Gly		
	10					15					20						
ccc	tgg	ttc	gtg	ccg	aag	gga	ссс	aac	cgc	gga	gtg	atc	atc	acc	atg		208
Pro	Trp	Phe	Val	Pro	Lys	Gly	Pro	Asn	Arg	Gly	Val	Ile	Ile	Thr	Met		
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ctg	gtc	gcc	acc	gcc	gtc	tgc	tgt	tac	ctc	ttc	tgg	ctc	atc	gcc	atc		256
Leu	Val	Ala	Thr	Ala	Val	Cys	Cys	Tyr	Leu	Phe	Trp	Leu	Ile	Ala	Ile		
				45				•	50					55			
ctg	gcg	cag	ctg	aac	ccc	ctg	ttc	ggg	ccc	cag	ctg	aag	aat	gag	acc		304
Leu	Ala	Gln	Leu	Asn	Pro	Leu	Phe	Gly	Pro	Gln	Leu	Lys	Asn	Glu	Thr		
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atc	tgg	tac	gtg	cgc	ttc	ctg	tgg	gag	tga	cccg	c¢ g	cccc	cgac	С			350
Ile	Trp	Tyr	Val	Arg	Phe	Leu	Trp	Glu									
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accatcctgc	tgggaactgg	gggggcctct	attgggttat	aggcaaggcc	ttttctctgg	830
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<210> 53

⟨211⟩ 1758

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

〈222〉 (190)... (1653)

<400> 53

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ggagg	gaaa	c at	g gg	g ga	t ac	t gg	c ct	g ag	a aa	g cg	g ag	a ga	g ga	t ga	g	228
		Ме	t Gl	y As	p Th	r Gl	y Le	u Ar	g Ly	s Ar	g Ar	g Gl	u As	p G1	u	
			1			;	5				1	0				
aag 1	tcg	atc	cag	agc	caa	gag	cct	aag	acc	acc	agt	ctc	caa	aag	gag	276
Lys S	Ser	Ile	Gln	Ser	Gln	Glu	Pro	Lys	Thr	Thr	Ser	Leu	Gln	Lys	Glu	
	15					20					25					
ctg	ggç	ctc	atc	agt	ggc	atc	tcc	atc	atc	gtg	ggc	acc	atc	att	ggc	324
Leu (Gly	Leu	Ile	Ser	Gly	Ile	Ser	Ile	Ile	Val	Gly	Thr	Ile	Ile	Gly	
30	•				35					40					45	
tct	ggg	atc	ttc	gtt	tcc	ccc	aag	tct	gtg	ctc	agc	aac	acg	gaa	gct	372
Ser	Gly	Ile	Phe	Val	Ser	Pro	Lys	Ser	Val	Leu	Ser	Asn	Thr	Glu	Ala	
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gtg	ggg	ccc	tgc	ctc	atc	ata	tgg	gcg	gct	tgc	ggg	gtc	ctc	gcg	acg	420
Val	Gly	Pro	Cys	Leu	Ile	Ile	Trp	Ala	Ala	Cys	Gly	Val	Leu	Ala	Thr	
			65					70					75			
ctg	ggt	gcc	ctg	tgc	ttt	gcg	gag	ctt	ggc	aca	atg	atc	acc	aag	tca	468
Leu	Gly	Ala	Leu	Cys	Phe	Ala	Glu	Leu	Gly	Thr	Met	Ile	Thr	Lys	Ser	
		80	ı				85	;				90)			
ggg	gga	gag	tat	ccc	tac	ctg	atg	gag	gcc	tac	ggg	ccc	ato	ccc	gcc	516
Gly	Gly	Glu	Tyr	Pro	Tyr	Leu	Met	Glu	Ala	Tyr	Gly	Pro	Ile	Pro	Ala	
	95					100					105	;				
tac	ctc	tto	tcc	tgg	g gcc	ago	ctg	g ato	gto	att	: aag	ccc	ac ₈	g tco	ttc	564

Tyr	Leu	rne	5er	irp	Ala	5er	Leu	116	vai	TIE	Lys	FIO	HILL	Ser	Lue	
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Ala	Ile	Ile	Cys	Leu	Ser	Phe	Ser	Glu	Tyr	Val	Cys	Ala	Pro	Phe	Tyr	
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Val	Gly	Cys	Lys	Pro	Pro	Gln	Ile	Val	Val	Lys	Cys	Leu	Ala	Ala	Ala	
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gcc	atc	ttg	ttc	atc	tcg	aca	gtg	aac	tca	ctg	agc	gtg	cgg	ctg	gga	708
Ala	Ile	Leu	Phe	Ile	Ser	Thr	Val	Asn	Ser	Leu	Ser	Val	Arg	Leu	Gly	
		160					165					170				
agc	tac	gtc	cag	aac	atc	ttc	acc	gcg	gcc	aag	ctg	gtg	atc	gtg	gcc	756
Ser	Tyr	Val	Gln	Asn	Ile	Phe	Thr	Ala	Ala	Lys	Leu	Val	Ile	Val	Ala	
	175					180					185					
atc	atc	atc	atc	agc	ggg	ctg	gtg	ctc	ctg	gcc	caa	gga	aac	aca	aag	804
Ile	Ile	Ile	Ile	Ser	Gly	Leu	Val	Leu	Leu	Ala	Gln	Gly	Asn	Thr	Lys	
190					195					200					205	
aat	ttt	gat	aat	tct	ttc	gag	ggc	gcc	cag	ctg	tct	gtg	gga	gcc	atc	852
Asn	Phe	Asp	Asn	Ser	Phe	Glu	Gly	Ala	Gln	Leu	Ser	Val	Gly	Ala	Ile	
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Ser	Leu	Ala	Phe	Tyr	Asn	Gly	Leu	Trp	Ala	Tyr	Asp	Gly	Trp	Asn	Gln	
			225					230					235	i		
ctc	aat	tac	atc	aca	gaa	gaa	ctt	aga	aac	cct	tac	aga	aac	ctg	cct	948
Leu	Asn	Tur	· Ile	Thr	- Glu	Glu	Leu	Arg	Asn	Pro	Tvr	Arg	Asn	Leu	Pro	

		240					245					250				
ttg	gcc	att	atc	atc	ggg	atc	ccc	ctg	gtg	acg	gcg	tgc	tac	atc	ctc	996
Leu	Ala	Ile	lle	Ile	Gly	Ile	Pro	Leu	Val	Thr	Ala	Cys	Tyr	Ile	Leu	
	255					260					265					
atg	aac	gtg	tcc	tac	ttc	acc	gtg	atg	act	gcc	acc	gaa	ctc	ctg	cag	1044
Met	Asn	Val	Ser	Tyr	Phe	Thr	Val	Met	Thr	Ala	Thr	Glu	Leu	Leu	Gln	
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tcc	cag	gcg	gtg	gct	gtg	aca	ttt	ggt	gac	cgt	gtt	ctc	tat	cct	gct	1092
Ser	Gln	Ala	Val	Ala	Val	Thr	Phe	Gly	Asp	Arg	Val	Leu	Tyr	Pro	Ala	
				290					295					300		
tct	tgg	atc	gtt	cca	ctt	ttt	gtg	gca	ttt	tca	acc	atc	ggt	gct	gct	1140
Ser	Trp	Ile	Val	Pro	Leu	Phe	Val	Ala	Phe	Ser	Thr	Ile	Gly	Ala	Ala	·
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aac	ggg	acc	tgc	ttc	aca	gcg	ggc	aga	ctc	att	tac	gtg	gcg	ggc	cgg	1188
Asn	Gly	Thr	Cys	Phe	Thr	Ala	Gly	Arg	Leu	Ile	Tyr	Val	Ala	Gly	Arg	
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gag	ggt	cac	atg	ctc	aaa	gtg	ctt	tct	tac	atc	agc	gtc	agg	cgc	ctc	1236
Glu	Gly	His	Met	Leu	Lys	Val	Leu	Ser	Tyr	Ile	Ser	Val	Arg	Arg	Leu	
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Thr	Pro	Ala	Pro	Ala	Ile	Ile	Phe	Tyr	Gly	Ile	Ile	Ala	Thr	Ile	Tyr	
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Ile	Ile	Pro	Gly	Asp	Ile	Asn	Ser	Leu	Val	Asn	Tyr	Phe	Ser	Phe	Ala	
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Phe	Thr	Arg	Lys	Glu	Leu	Glu	Arg	Pro	Ile	Lys	Val	Pro	Val	Val	Ile	
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ccc	gtc	ttg	atg	aca	ctc	atc	tct	gtg	ttt	ttg	ġtt	ctg	gct	cca	atc	1476
Pro	Val	Leu	Met	Thr	Leu	Ile	Ser	Val	Phe	Leu	Val	Leu	Ala	Pro	Ile	
	415					420					425					
atc	agc	aag	ссс	acc	tgg	gag	tac	ctc	tac	tgt	gtg	ctg	ttt	ata	tta	. 1524
Ile	Ser	Lys	Pro	Thr	Trp	Glu	Tyr	Leu	Tyr	Cys	Val	Leu	Phe	Ile	Leu	
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Ser	Gly	Leu	Leu	Phe	Tyr	Phe	Leu	Phe	Val	His	Tyr	Lys	Phe	Gly	Trp	
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gct	cag	aaa	atc	tca	aag	ccg	att	acc	atg	cac	ctt	cag	atg	cta	atg	1620
Ala	Gln	Lys	Ile	Ser	Lys	Pro	Ile	Thr	Met	His	Leu	Gln	Met	Leu	Met	
			465					470					475			
gaa	gtg	gtc	cca	ccg	gag	gaa	gac	cct	gag	taad	caago	ctc (cgtc	tctt	gt	1670
Glu	Val	Val	Pro	Pro	Glu	Glu	Asp	Pro	Glu							
		480					485									
agc	caagi	tca (gctga	aatt	ta t	tttc	ttaa	g ca	atati	ttgt	ggt	tatt	tct ·	tcct	ttttt	1730
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Met Thr Pro Gln Pro Ala Gly	
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ccc ccg gat ggg ggc tgg ggc tgg gtg gtg gcg gcc gca gcc ttc gcg	222
Pro Pro Asp Gly Gly Trp Gly Trp Val Val Ala Ala Ala Ala Phe Ala	
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ata aac ggg ctg tcc tac ggg ctg ctg cgc tcg ctg ggc ctt gcc ttc	270
Ile Asn Gly Leu Ser Tyr Gly Leu Leu Arg Ser Leu Gly Leu Ala Phe	
25 30 35	
cct gac ctt gcc gag cac ttt gac cga agc gcc cag gac act gcg tgg	318
Pro Asp Leu Ala Glu His Phe Asp Arg Ser Ala Gln Asp Thr Ala Trp	
40 45 50 55	
atc agc gcc ctg gcc ctg gcc gtg cag cag gca gcc agc ccc gtg ggc	366
Ile Ser Ala Leu Ala Leu Ala Val Gln Gln Ala Ala Ser Pro Val Gly	
60 65 70	
age gee etg age acg ege tgg ggg gee ege eee gtg gtg atg gtt ggg	414

Ser	Ala	Leu	Ser	Thr	Arg	Trp	Gly	Ala	Arg	Pro	Val	Val	Met	Val	Gly	
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ggc	gtc	ctc	gcc	tcg	ctg	ggc	ttc	gtc	ttc	tcg	gct	ttc	gcc	agc	ggt	462
Gly	Val	Leų	Ala	Ser	Leu	Gly	Phe	Val	Phe	Ser	Ala	Phe	Ala	Ser	Gly	
		90					95					100				
ctg	ctg	cat	ctc	tac	ctc	ggc	ctg	ggc	ctc	ctc	gct	ggc	ttt	ggt	tgg	510
Leu	Leu	His	Leu	Tyr	Leu	Gly	Leu	Gly	Leu	Leu	Ala	Gly	Phe	Gly	Trp	
	105					110					115					
gcc	ctg	gtg	ttc	gcc	ccc	gcc	cta	ggc	acc	ctc	tcg	cgt	tac	ttc	tcc	558
Ala	Leu	Val	Phe	Ala	Pro	Ala	Leu	Gly	Thr	Leu	Ser	Arg	Tyr	Phe	Ser	
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Arg	Arg	Arg	Val	Leu	Ala	Val	Gly	Leu	Ala	Leu	Thr	Gly	Asn	Gly	Ala	
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tcc	tcg	ctg	ctc	ctg	gcg	ccc	gcc	ttg	cag	ctt	ctc	ctc	gat	act	ttc	654
Ser	Ser	Leu	Leu	Leu	Ala	Pro	Ala	Leu	Gln	Leu	Leu	Leu	Asp	Thr	Phe	
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ggc	tgg	cgg	ggc	gct	ctg	ctc	ctc	ctc	ggc	gcg	atc	acc	ctc	cac	ctc	702
G1y	Trp	Arg	Gly	Ala	Leu	Leu	Leu	Leu	Gly	Ala	Ile	Thr	Leu	His	Leu	
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acc	ccc	tgt	ggc	gcc	ctg	ctg	cta	ccc	ctg	gtc	ctt	cct	gga	gac	ccc	750
Thr	Pro	Cys	Gly	Ala	Leu	Leu	Leu	Pro	Leu	Val	Leu	Pro	Gly	Asp	Pro	
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cca	gcc	cca	ccg	cgt	agt	ccc	cta	gct	gcc	ctc	ggc	ctg	agt	ctg	ttc	798
Pro	Ala	Pro	Pro	Arg	Ser	Pro	Leu	Ala	Ala	Leu	Gly	Leu	Ser	Leu	Phe	

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Thr	Arg	Arg	Ala	Phe	Ser	Ile	Phe	Ala	Leu	Gly	Thr	Ala	Leu	Val	Gly	
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Gly	Gly	Tyr	Phe	Val	Pro	Tyr	Val	His	Leu	Ala	Pro	Arg	Phe	Arg	Pro	
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Gly	Pro	Gly	Gly	Ile	Arg	Ser	Ser	Ala	Gly	Gly	Gly	Arg	Gly	Cys	Asp	
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Gly	Gly	Cys	Gly	Arg	Pro	Ala	Gly	Leu	Arg	Val	Ala	Gly	Arg	Pro	Arg	. . ,
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Leu	Gly	Ala	Pro	Pro	Ala	Ala	Ala	Gly	Arg	Ile	Arg	Gly	Ser	Asp	Trp	
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gct	ggg	gct	gtg	ggt	ggt	ggg	gct	ggt	gcc	cgt	ggt	ggg	cgg	cga	aga	1086
Ala	Gly	Ala	Val	Gly	Gly	Gly	Ala	Gly	Ala	Arg	Gly	Gly	Arg	Arg	Arg	
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gag	ctg	ggg	ggg	tcc	cct	gct	ggc	cgc	ggc	tgt	ggc	cta	tgg	gct	gag	1134
Glu	Leu	Gly	Gly	Ser	Pro	Ala	Gly	Arg	Gly	Cys	Gly	Leu	Trp	Ala	Glu	
			315					320					325			
cgc	ggg	gag	tta	cgc	ccc	gct	ggt	ttt	cgg	tgt	act	ccc	cgg	gct	ggt	1182
Arg	Gly	Glu	Leu	Arg	Pro	Ala	Gly	Phe	Arg	Cys	Thr	Pro	Arg	Ala	Gly	
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ggg cgt cgg agg tgt	ggt gca ggc cac	agg gct ggt gat ga	at gct gat 1230
Gly Arg Arg Cys	Gly Ala Gly His	Arg Ala Gly Asp As	sp Ala Asp
345	350	355	
gag cct cgg ggg gct	cct ggg ccc tcc	cct gtc agg ctt co	ct aag gga 1278
Glu Pro Arg Gly Ala	Pro Gly Pro Ser	Pro Val Arg Leu Pa	ro Lys Gly
360	365	370	375
tg agacaggaga cttca	ccgcc tctttcctcc	tgtctggttc tttgatc	cete 1330
tccggcagct tcatctac	at agggttgccc agg	ggcgctgc cctcctgtgg	g tecageetee 1390
cctccagcca cgcctccc	cc agagacgggg gag	getgette cegeteces	a ggcagtcttg 1450
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⟨220⟩

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Met Ala Thr Thr Ala

5

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gcg ccg gcg ggc gcc cga aat gga gct ggc ccg gaa tgg gga ggg 163

Ala	Pro	Ala	Gly	Gly	Ala	Arg	Asn	Gly	Ala	Gly	Pro	Glu	Trp	Gly	Gly	
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Phe	Glu	Glu	Asn	Ile	Gln	Gly	Gly	Gly	Ser	Ala	Val	Ile	Asp	Met	Glu	
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Asn	Met	Asp	Asp	Thr	Ser	Gly	Ser	Ser	Phe	Glu	Asp	Met	Gly	Glu	Leu	
		40					45					50				
cat	cag	cgc	ctg	cgc	gag	gaa	gaa	gta	gac	gct	gat	gca	gct	gat	gca	307
His	Gln	Arg	Leu	Arg	Glu	Glu	Glu	Val	Asp	Ala	Asp	Ala	Ala	Asp	Ala	
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gct	gct	gct	gaa	gag	gag	gat	gga	gag	ttc	ctg	ggc	atg	aag	ggc	ttt	355
Ala	Ala	Ala	Glu	Glu	Glu	Asp	Gly	Glu	Phe	Leu	Gly	Met	Lys	Gly	Phe	
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Lys	Gly	Gln	Leu	Ser	Arg	Gln	Val	Ala	Asp	Gln	Met	Trp	Gln	Ala	Gly	
				90					95					100		
		caa														451
Lys	Arg	Gln	Ala	Ser	Arg	Ala	Phe	Ser	Leu	Tyr	Ala	. Asn			Ile	
			105					110					115	i		
ctc	aga	ccc	tac	ttt	gat	gtg	gag	cct	gct	cag	gtg	cga	ago	agg	ctc	499
Leu	Arg	Pro	Tyr	Phe	Asp	Val	Glu	Pro	Ala	Gln	Val	Arg	Ser	· Arg	g Leu	
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1	C1.		. Mo+	- 11-	Pro	. 112	100	Mo+	Vo1	Acr	Phe	Pro	Gli	Lvs	: Ile	

	135					140					145					
gca	ggt	gaa	ctc	tat	gga	cct	ctc	atg	ctg	gtc	ttc	act	ctg	gtt	gct	595
Ala	Gly	Glu	Leu	Tyr	Gly	Pro	Leu	Met	Leu	Val	Phe	Thr	Leu	Val	Ala	
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atc	cta	ctc	cat	ggg	atg	aag	acg	tct	gac	act	att	atc	cgg	gag	ggc	643
Ile	Leu	Leu	His	Gly	Met	Lys	Thr	Ser	Asp	Thr	Ile	Ile	Arg	Glu	Gly	
				170					175					180		
acc	ctg	atg	ggc	aca	gcc	att	ggc	acc	tgc	ttc	ggc	tac	tgg	ctg	gga	691
Thr	Leu	Met	Gly	Thr	Ala	Ile	Gly	Thr	Cys	Phe	Gly	Tyr	Trp	Leu	Gly	
			185					190					195			
gtc	tca	tcc	ttc	att	tac	ttc	ctt	gcc	tac	ctg	tgc	aac	gcc	cag	atc	739
Val	Ser	Ser	Phe	Ile	Tyr	Phe	Leu	Ala	Tyr	Ĺeu	Cys	Asn	Ala	Gln	Ile	
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Thr	Met	Leu	Gln	Met	Leu	Ala	Leu	Leu	Gly	Tyr	Gly	Leu	Phe	Gly	His	
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Cys	Ile	Val	Leu	Phe	Ile	Thr	Tyr	Asn	Ile	His	Leu	His	Ala	Leu	Phe	
230					235					240	•				245	
tac	cto	tto	tgg	ctg	ttg	gtg	ggt	gga	ctg	tcc	aca	ctg	cgo	atg	gta	883
Tyr	Leu	Phe	Trp	Leu	Leu	Val	Gly	Gly	Leu	Ser	Thr	Leu	Arg	g Met	Val	
				250)				255	;				260)	
gca	gte	g ttg	ggtg	tct	cgg	acc	gtg	ggg	ccc	aca	cag	cgg	ct	g ctc	ctc	93
Ala	Va]	Leu	ı Val	Ser	Arg	Thr	· Val	l Gly	Pro	Thr	Glr	Arg	Le	u Leu	ı Leu	
			265	5				270)				27	5		

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Phe	Ala	Tyr	His	Lys	Val	Val	Glu	Gly	Ile	Leu	Asp	Thr	Leu	Glu	Gly	
	295					300					305					
ccc	aac	atc	ccg	ccc	atc	cag	agg	gtc	ccc	aga	gac	atc	cct	gcc	atg	1075
Pro	Asn	Ile	Pro	Pro	Ile	Gln	Arg	Val	Pro	Arg	Asp	Ile	Pro	Ala	Met	
310					315					320					325	
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Leu	Pro	Ala	Ala	Arg	Leu	Pro	Thr	Thr	Val	Leu	Asn	Ala	Thr	Ala	Lys	
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gct	gtt	gcg	gtg	acc	ctg	cag	tca	cac	tga	cccc	acc '	tgaa:	attc	tt		1170
			Val						Ū			•				
7110	,,,,	1110	345		200	011.	001	350								
						- 4					+-+	+	~~^	oogt.	ootgot	1230
															cctgat	
gac	atgt	ttc	gtag	atgg	gg t	ttgc	agct	g cc	actg	agct	gta	gctg	cgt	aagt	acctcc	1290
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ctt	cctt	cct	cttt	atct	ct c	ccac	attg	t ct	tgct	aaat	ata	gact	tgg	taat	taaaat	1470
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					Met	Ser	Ser	Gln	Pro) Ala	Gly	Asr	Glr	n Thi	Ser	
					1				5	5				10)	
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Pro	Gly	Ala	Thr	Glu	Asp	Tyr	Ser	Tyr	Gly	Ser	Trp	Tyr	Ile	Asp	Glu	
		•	15					20					25	•		
ccc	cag	ggg	ggc	gag	gag	ctc	cag	cca	gag	ggg	gaa	gtg	ссс	tcc	tgc	208
Pro	Gln	Gly	Gly	Glu	Glu	Leu	Gln	Pro	Glu	Gly	Glu	Val	Pro	Ser	Cys	
		30					35					40				
cac	acc	agc	ata	cca	ccc	ggc	ctg	tac	cac	gcc	tgc	ctg	gcc	tcg	ctg	256
His	Thr	Ser	Ile	Pro	Pro	Gly	Leu	Tyr	His	Ala	Cys	Leu	Ala	Ser	Leu	
	45					50					55					
tca	atc	ctt	gtg	ctg	ctg	ctc	ctg	gcc	atg	ctg	gtg	agg	cgc	cgc	cag	304
Ser	Ile	Leu	Val	Leu	Leu	Leu	Leu	Ala	Met	Leu	Val	Arg	Arg	Arg	Gln	
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ctc	tgg	cct	gac	tgt	gtg	cgt	ggc	agg	ccc	ggc	ctg	ccc	agc	cct	gtg	352
Leu	Trp	Pro	Asp	Cys	Val	Arg	Gly	Arg	Pro	Gly	Leu	Pro	Ser	Pro	Val	
				80					85	;				90	ı	
gat	ttc	ttg	gct	ggg	gac	agg	ccc	cgg	gca	gtg	cct	gct	gct	gtt	ttc	400

Asp	Phe	Leu	Ala	Gly	Asp	Arg	Pro	Arg	Ala	Val	Pro	Ala	Ala	Val	Phe	
			95					100					105			
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Met	Val	Leu	Leu	Ser	Ser	Leu	Cys	Leu	Leu	Leu	Pro	Asp	Glu	Asp	Ala	
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Leu	Pro	Phe	Leu	Thr	Leu	Ala	Ser	Ala	Pro	Ser	Gln	Asp	Gly	Lys	Thr	
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gag	gct	cca	aga	ggg	gcc	tgg	aag	ata	ctg	gga	ctg	ttc	tat	tat	gct	544
Glu	Ala	Pro	Arg	Gly	Ala	Trp	Lys	Ile	Leu	Gly	Leu	Phe	Tyr	Tyr	Ala	
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gcc	ctc	tac	tac	cct	ctg	gct	gcc	tgt	gcc	acg	gct	ggc	cac	aca	gct	592
Ala	Leu	Tyr	Tyr	Pro	Leu	Ala	Ala	Cys	Ala	Thr	Ala	G1y	His	Thr	Ala	
				160)				165					170		
gca	cac	ctg	cto	ggc	agc	acg	ctg	tcc	tgg	gcc	cac	ctt	ggg	gtc	cag	640
Ala	His	. Leu	ı Leu	Gly	Ser	Thr	Leu	Ser	Trp	Ala	His	Leu	Gly	Val	Gln	
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gto	tg:	g cas	g agg	g gca	a gag	tgt	ccc	cag	gtg	ccc	888	ato	tac	aag	tac	688
Va]	l Tr	Glı	n Arg	g Ala	ı Glu	ı Cys	Pro	Glr	Val	Pro	Lys	Ile	Tyr	Lys	Tyr	
		190	0				195	5				200)			
tac	c tc	c ct	g cta	g gco	c to	ctg	cct	cto	cte	g ctg	ggo	cto	gga	tto	ctg	736
Ty	r Se	r Le	u Lei	u Ala	a Ser	r Lei	ı Pro	Leu	ı Let	ı Lev	ı Gly	, Le	ı Gly	7 Pho	e Leu	
	20	5				210)				215	5				
ag	c ct	t tg	g ta	c cc	t gt	g cag	g ct	g gti	g aga	a ago	tt:	c ag	c cg ¹	t ag	g aca	784
															g Thr	

220					225					230					235	
gga	gca	ggc	tcc	aag	ggg	ctg	cag	agc	agc	tac	tct	gag	gaa	tat	ctg	. 832
Gly	Ala	Gly	Ser	Lys	Gly	Leu	Gln	Ser	Ser	Tyr	Ser	Glu	Glu	Tyr	Leu	
				240					245					250		
agg	aac	ctc	ctt	tgc	agg	aag	aag	ctg	gga	agc	agc	tac	cac	acc	tcc	880
Arg	Asn	Leu	Leu	Cys	Arg	Lys	Lys	Leu	Gly	Ser	Ser	Tyr	His	Thr	Ser	
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Lys	His	Gly	Phe	Leu	Ser	Trp	Ala	Arg	Val	Cys	Leu	Arg	His	Cys	Ile	
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Tyr	Thr	Pro	Gln	Pro	Gly	Phe	His	Leu	Pro	Leu	Lys	Leu	Val	Leu	Ser	
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Ala	Thr	Leu	Thr	Gly	Thr	Ala	Ile	Tyr	Gln	Val	Ala	Leu	Leu	Leu	Leu	
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gtg	ggc	gtg	gta	ccc	act	atc	cag	aag	gtg	agg	gca	ggg	gtc	acc	acg	1072
Val	G1y	Val	Val	Pro	Thr	Ile	Gln	Lys	Val	Arg	Ala	Gly	Val	Thr	Thr	
				320		•			325					330		
gat	gtc	tcc	tac	ctg	ctg	gcc	ggc	ttt	gga	atc	gtg	ctc	tcc	gag	gac	1120
Asp	Val	Ser	Tyr	Leu	Leu	Ala	Gly	Phe	Gly	Ile	Val	Leu	Ser	Glu	Asp	
			335					340					345			
aag	cag	gag	gtg	gtg	gag	ctg	gtg	aag	cac	cat	ctg	tgg	gct	ctg	gaa	1168
Lys	Gln	Glu	Val	Val	Glu	Leu	Val	Lys	His	His	Leu	Trp	Ala	Leu	Glu	
		350					355					360				

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Val	Cys	Tyr	Ile	Ser	Ala	Leu	Val	Leu	Ser	Cys	Leu	Leu	Thr.	Phe	Leu	•
	365		٠			370					375				•	
gtc	ctg	atg	cgc	tca	ctg	gtg	aca	cac	agg	acc	aac	ctt	cga	gct	ctg	1264
Val	Leu	Met	Arg	Ser	Leu	Val	Thr	His	Arg	Thr	Asn	Leu	Arg	Ala	Leu	
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cac	cga	gga	gct	gcc	ctg	gac	ttg	agt	ccc	ttg	cat	cgg	agt	ccc	cat	1312
His	Arg	Gly	Ala	Ala	Leu	Asp	Leu	Ser	Pro	Leu	His	Arg	Ser	Pro	His	
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ccc	tcc	cgc	caa	gcc	ata	ttc	tgt	tgg	atg	agc	ttc	agt	gcc	tac	cag	1360
Pro	Ser	Arg	Gln	Ala	Ile	Phe	Cys	Trp	Met	Ser	Phe	Ser	Ala	Tyr	Gln	
			415					420					425			
aca	gcc	ttt	atc	tgc	ctt	ggg	ctc	ctg	gtg	cag	cag	atc	atc	ttc	ttc	1408
Thr	Ala	Phe	Ile	Cys	Leu	Gly	Leu	Leu	Val	Gln	Gln	Ile	Ile	Phe	Phe	
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ctg	gga	acc	acg	gcc	ctg	gcc	ttc	ctg	gtg	ctc	atg	cct	gtg	ctc	cat	1456
Leu	Gly	Thr	Thr	Ala	Leu	Ala	Phe	Leu	Val	Leu	Met	Pro	Val	Leu	His	
	445					450	1				455					
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Gly	Arg	Asr	Leu	Leu	Leu	Phe	Arg	Ser	Leu	Glu	Ser	Ser	Trp	Pro	Phe	
460)				465	,				470)				475	
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Trp	Leu	ı Thi	r Leu	Ala	Leu	ı Ala	val	Ile			Asr	Met	. Ala		His	
				480					485					490		
tgg	ggto	tte	cte	g gag	g act	t cat	t gat	gga	a cac	cce	cas	g ctg	gaco	88	cgg	1600

Trp	Val	Phe	Leu	Glu	Thr	His	Asp	Gly	His	Pro	Gln	Leu	Thr	Asn	Arg	
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Arg	Val	Leu	Tyr	Ala	Ala	Thr	Phe	Leu	Leu	Phe	Pro	Leu	Asn	Val	Leu	
		510					515					520				
gtg	ggt	gcc	atg	gtg	gcc	acc	tgg	cga	gtg	ctc	ctc	tct	gcc	ctc	tac	1696
Val	Gly	Ala	Met	Val	Ala	Thr	Trp	Arg	Val	Leu	Leu	Ser	Ala	Leu	Tyr	
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aac	gcc	atc	cac	ctt	ggc	cag	atg	gac	ctc	agc	ctg	ctg	cca	ccg	aga	1744
Asn	Ala	Ile	His	Leu	Gly	Gln	Met	Asp	Leu	Ser	Leu	Leu	Pro	Pro	Arg	
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gcc	gcc	act	ctc	gac	ccc	ggc	tac	tac	acg	tac	cga	aac	ttc	ttg	aag	1792
Ala	Ala	Thr	Leu	Asp	Pro	Gly	Tyr	Tyr	Thr	Tyr	Arg	Asn	Phe	Leu	Lys	
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Ile	Glu	Val	Ser	Gln	Ser	His	Pro	Ala	Met	Thr	Ala	Phe	Cys	Ser	Leu	
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ctc	ctg	caa	gcg	cag	agc	ctc	cta	ccc	agg	acc	atg	gca	gcc	ccc	cag	1888
Leu	Leu	Gln	Ala	Gln	Ser	Leu	Leu	Pro	Arg	Thr	Met	Ala	Ala	Pro	Gln	
		590					595					600				
gac	agc	ctc	aga	cca	ggg	gag	gaa	gac	gaa	ggg	atg	cag	ctg	cta	cag	1936
Asp	Ser	Leu	Arg	Pro	Gly	Glu	Glu	Asp	Glu	Gly	Met	Gln	Leu	Leu	Gln	
	605					610					615	;				
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Thr	Lys	Asp	Ser	Met	Ala	Lys	Gly	Ala	Arg	Pro	Gly	Ala	Ser	Arg	Gly	

121/307

620	62	5	630	635	
agg gct cg	c tgg ggt ct	g gcc tac a	cg ctg ctg cac	aac cca acc ctg	2032
Arg Ala Ar	g Trp Gly Le	u Ala Tyr T	hr Leu Leu His	Asn Pro Thr Leu	
	640		645	650	
cag gtc tt	c cgc aag ac	g gcc ctg t	tg ggt gcc aat	ggt gcc cag ccc	2080
Gln Val Ph	e Arg Lys Th	ır Ala Leu L	eu Gly Ala Asn	Gly Ala Gln Pro	
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⟨220⟩

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122/307

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Met Ile Val Cys Leu Leu Phe Met Ile	
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Leu Leu Ala Lys Glu Val Gln Leu Val Asp Gln Thr Asp Ser Pro Leu	
15 20 25	
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Leu Ser Leu Leu Gly Gln Thr Ser Ser Leu Ser Trp His Leu Val Asp	
30 35 40	
att gtg tcg tac cag agt gtg cta agt tat ttc agc agc cat tac ccg	256
Ile Val Ser Tyr Gln Ser Val Leu Ser Tyr Phe Ser Ser His Tyr Pro	
45 50 55	
ccg tcc atc atc ctg gca aaa gaa tct tat gct gaa tta atc atg aag	304
Pro Ser Ile Ile Leu Ala Lys Glu Ser Tyr Ala Glu Leu Ile Met Lys	
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ctc cta aaa gtg tct gcg ggc ctt tct att cct act gac agc cag aag	352
Leu Leu Lys Val Ser Ala Gly Leu Ser Ile Pro Thr Asp Ser Gln Lys	
75 80 85 90	
cat ctt gat gca gtt cca aaa tgc caa gct ttt act cat cag atg gtt	400
His Leu Asp Ala Val Pro Lys Cys Gln Ala Phe Thr His Gln Met Val	

100

caa ttc ctc agc acc ctg gaa caa aat gga aaa atc acc tta gca gtc

95

105

448

Gln	Phe	Leu	Ser	Thr	Leu	Glu	Gln	Asn	Gly	Lys	Ile	Thr	Leu	Ala	Val	
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cta	gaa	cag	gaa	atg	tct	aag	ctc	tta	gac	gat	atc	att	gtc	ttt	aac	496-
Leu	Glu	Gln	Glu	Met	Ser	Lys	Leu	Leu	Asp	Asp	Ile	Ile	Val	Phe	Asn	
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ccg	ccc	gac	atg	gac	agc	cag	acc	cgc	cac	atg	gcc	ctc	agc	agc	ctc	544
Pro	Pro	Asp	Met	Asp	Ser	Gln	Thr	Arg	His	Met	Ala	Leu	Ser	Ser	Leu	
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Phe	Met	Glu	Val	Leu	Met	Met	Met	Asn	Asn	Ala	Thr	Ile	Pro	Thr	Ala	
155					160		٠			165					170	
gag	ttc	ctt	cgg	ggc	agt	atc	cgg	acc	tgg	att	ggc	caa	aaa	atg	cat	640
Glu	Phe	Leu	Arg	Gly	Ser	Ile	Arg	Thr	Trp	Ile	Gly	Gln	Lys	Met	His	
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ggg	ctg	gtg	gtg	ctg	ccc	ctt	tta	aca	gca	gcc	tgc	cag	gago	ctg	gcg	688
Gly	Leu	Val	Val	Leu	Pro	Leu	Leu	Thr	Ala	Ala	Cys	Glr	s Ser	Leu	Ala	
			190	ı				195	;				200)		
tcc	gto	cgo	cac	atg	gct	gag	act	aca	gaa	gcc	tgc	ato	act	gcc	tac	736
Ser	· Va]	Arg	g His	Met	. Ala	Glu	Thr	Thr	Glu	Ala	Cys	Ile	e Thù	r Ala	Tyr	
		205	5				210	,				215	5			
tto	aaa	a gaa	a ago	cct	cto	aat	cag	aat	t tca	gga	tgg	gg	a cc	c ati	tctg	784
Phe	. Lys	s Glu	ı Ser	Pro) Leu	ı Aşr	Gln	Ası	ı Ser	Gly	Trp	G1	y Pr	o Ile	e Leu	
	220)				225	5				230)				
gta	a tc	c ct	t ca	g gti	t ccc	gag	g ctc	ace	c atg	gaa	gag	tt	c ct	g ca	g gag	832
Va	می ا	r i a	C1,	n Va`	l Pro	s Glu	ı Lei	ı Th	r Met	: G1:	ı Glı	ı Ph	e Le	u Gl:	n Glu	

2	235					240					245					250	
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(Cys	Leu	Thr	Leu	Gly	Ser	Tyr	Leu	Thr	Leu	Tyr	Val	Tyr	Leu	Leu	Gln	
					255					260					265		
	tgt	tta	aac	agc	gaa	cag	act	tta	agg	aat	gaa	atg	aaa	gtg	ctg	ctc	928
(Cys	Leu	Asn	Ser	Glu	Gln	Thr	Leu	Arg	Asn	Glu	Met	Lys	Val	Leu	Leu	
		•		270					275					280			
	atc	tta	agc	aag	tgg	ctg	gaa	cag	gtg	tac	cca	agc	tcc	gtg	gag	gaa	976
	Ile	Leu	Ser	Lys	Trp	Leu	G1u	Gln	Val	Tyr	Pro	Ser	Ser	Val	Glu	Glu	٠
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	Glu	Ala	Lys	Leu	Phe	Leu	Trp	Trp	His	Gln	Val	Leu	Gln	Leu	Ser	Leu	
		300					305					310					
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	Ile	Gln	Thr	Glu	Gln	Asn	Asp	Ser	Val	Leu	Thr	Glu	Ser	Val	Ile	Arg	
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	Ile	Leu	Leu	Leu	Val	Gln	Ser	Arg	Gln	Asn	Leu	Val	Ala	Glu	Glu	Arg	
					335	•				340	İ				345	5	
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	Leu	Ser	Ser	Gly	Ile	Leu	Gly	Ala	Ile	Gly	Phe	Gly	Arg	g Lys	S Sea	r Pro	
				350)				355	5				360)		
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	Leu	Sei	- Asr	n Arg	, Phe	e Arg	y Val	l Val	l Ala	Arg	g Ser	Me1	t Ala	a Ala	a Ph	e Leu	
			369	5				370)				379	5			

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Ser	Val	Gln	Val	Pro	Met	Glu	Asp	Gln	Ile	Arg	Leu	Arg	Pro	Gly	Ser	
•	380					385				•	390	•			•	•
gaa	tta	cat	ctg	acc	ссс	aaa	gct	cag	cag	gct	ctg	aat	gct	ctt	gaa	1312
Glu	Leu	His	Leu	Thr	Pro	Lys	Ala	Gln	Gln	Ala	Leu	Asn	Ala	Leu	Glu	
395					400					405					410	
tcc	atg	gca	tca	agt	aag	cag	tat	gtt	gaa	tac	cag	gat	caa	ata	ttg	1360
Ser	Met	Ala	Ser	Ser	Lys	Gln	Tyr	Val	Glu	Tyr	Gln	Asp	Gln	Ile	Leu	
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caa	gcc	acc	caa	ttt	ata	agg	cat	cct	ggc	cat	tgc	ctt	caa	gat	ggg	1408
Gln	Ala	Thr	Gln	Phe	Ile	Arg	His	Pro	Gly	His	Cys	Leu	Gln	Asp	Gly	
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Lys	Ser	Phe	Leu	Ala	Leu	Leu	Val	Asn	Cys	Leu	Tyr	Pro	Glu	Val	His	
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Tyr	Leu	Asp	His	Ile	Arg											
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gta	tgtti	tcc a	actto	ctgt	ct ci	gtti	ttatį	g taa	aatgi	ttcc	agat	tctga	aca a	accti	tggaag	1870
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Cys	His	Leu	Trp	Tyr	Ala	Val	Leu	Cys	Gly	Gln	Leu	Ala	Glu	His	Glu	
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Lys	Asr	n Ser	Glu	ı Ala	Are	Trp	Trp	Met	t Lys	Leu	Ala	ı Lei	ı Glu	ı Leu	Pro	
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Ası	Va.	1 Thi	r Ly:	s Glu	ı Ası) Let	ı Ala	a Ile	e Glr	ı Lys	s Asj	p Lei	u Gli	u Glu	ı Leu	
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G1	u Va	1 I1	e Le	u Ar	g As	p										
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Met Glu Gln Gly Ser Gly Arg Leu Glu Asp Phe Pro

1 5 10

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157

Val Asn Val Phe Ser Val Thr Pro Tyr Thr Pro Ser Thr Ala Asp Ile

15 20 25

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		Me	t Ala Gly A	la Glu Aspí	Trp Pro Gly	
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Gln Gln L	eu Glu Leu	Asp Glu Asp	Glu Ala Se	r Cys Cys A	rg Trp Gly	
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209

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Ala	His	Asn	Leu	Val	His	Leu	Leu	Leu	Leu	Ala	Arg	Trp	Glu	Asp	Thr	
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Phe	Ar	g Thi	r His	Se Se	r Pro	Glu	ı Ala	a Lei	ı Glu	Glr	ı Lei	т Ту	r Pro	Tr	Glu	

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Cys	Phe	Val	Phe	Cys	Leu	Ile	Ile	Phe	Gly	Thr	Phe	Thr	Asn	Gln	Ile	
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cac	aag	tgg	tcg	cac	acg	tac	ttt	ggg	ctg	cca	cgc	tgg	gtc	acc	ctc	689
His	Lys	Trp	Ser	His	Thr	Tyr	Phe	Gly	Leu	Pro	Arg	Trp	Val	Thr	Leu	
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250)				255	;				260)				265	
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Ala	Glr	Lys	s Ile	e Lys	S											
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ggtttaccca	ctccctgcac	gcctgccgtc	cctgccccgc	tgggcagccc	ttcagtgtgg	1230
ctggcgttgg	ggccagtgag	ttgcctcttt	ccctccttgt	ctggccccag	tggtctgggg	1290
agcccccagg	cacacctaag	cgtcgtggag	cattgttctg	ccacagccct	gcatactgac	1350
cccgggaggc	tgggcaggtg	gacagcccca	gccaccacct	tcagcctagc	ctgtccccca	1410
aggatggtga	agctcagcag	gggtctgagg	gtagccggcc	agaagaggct	ggaacctcct	1470
gctcaagtct	agacccctac	ttctctgctg	ccccaccct	gccagagctg	atgtttccaa	1530
taccaagatg	tcttcacagg	gcacagcccc	tgcagagcat	cttggtcatt	tggaagagga	1590
cacggtatcc	cctctggcca	gagtatgtca	gagaaggaag	agtagggctt	ttttgttttg	1650
tttttttta	aaggtgcttg	cttgtttaat	gtaaataata	gaaagcctta	atatcttttc	1710
tgtaacacgg	agtaatattt	taatgtcatg	ttttggatgt	acataatata	tttataacaa	1770
agcagcaaga	gtctactt					1788

<210> 61

<211> 389

<212> PRT

<213> Homo sapiens

<400> 61

Met Asp Arg Gly Glu Lys Ile Gln Leu Lys Arg Val Phe Gly Tyr Trp

1 5 10 15

Trp Gly Thr Ser Phe Leu Leu Ile Asn Ile Ile Gly Ala Gly Ile Phe
20 25 30

Val Ser Pro Lys Gly Val Leu Ala Tyr Ser Cys Met Asn Val Gly Val

35 40 45

Ser Leu Cys Val Trp Ala Gly Cys Ala Ile Leu Ala Met Thr Ser Thr

	50					55					60				
Leu	Cys	Ser	Ala	Glu	Ile	Ser	Ile	Ser	Phe	Pro	Cys	Ser	Gly	Ala	Gln
65					70					75					80
Tyr	Tyr	Phe	Leu	Lys	Arg	Tyr	Phe	Gly	Ser	Thr	Val	Ala	Phe	Leu	Asn
				85					90					95	
Leu	Trp	Thr	Ser	Leu	Phe	Leu	Gly	Ser	Gly	Val	Val	Ala	Gly	Gln	Ala
•			100					105					110		
Leu	Leu	Leu	Ala	Glu	Tyr	Ser	Ile	Gln	Pro	Phe	Phe	Pro	Ser	Cys	Ser
		115					120					125			
Val	Pro	Lys	Leu	Pro	Lys	Lys	Cys	Leu	Ala	Leu	Ala	Met	Leu	Trp	Ile
	130					135					140				
Val	Gly	Ile	Leu	Thr	Ser	Arg	Gly	Val	Lys	Glu	Val	Thr	Trp	Leu	Gln
145					150					155	i				160
Ile	Ala	Ser	Ser	· Val	Leu	Lys	Val	Ser	· Ile	Leu	Ser	Phe	lle	Ser	Leu
				165	,				170)				175	
Thr	Gly	Va]	[Va]	Phe	Leu	Ile	Arg	Gly	Lys	Lys	s Glu	Asr	ı Val	Glu	Arg
			180)				185	5				190	١	
Phe	G1r	n Ası	n Ala	a Phe	. Asp	Ala	ı Glı	ı Lev	ı Pro	o Ası	o Ile	e Sei	r His	Leu	Ile
		19	5				200)				209	5		
Gln	Ala	a Ile	e Pho	e Glr	1 Gly	, Tyj	r Phe	e Ala	a Ty	r Se	r Gly	y Glı	u Leu	Lys	Lys
	210	0				219	5				220	0			
Pro	Ar	g Th	r Th	r Ile	e Pro	o Ly:	s Cy:	s Il	e Ph	e Th	r Ala	a Le	u Pro	Leu	ı Val
225	5				23	0				23	5				240
Thr	- Va	l Va	l Ty	r Le	u Le	u Va	l As	n Il	e Se	r Ty	r Le	u Th	r Va	l Le	u Thi
				24	5				25	0				25	5

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Pro Arg Glu Ile Leu Ser Ser Asp Ala Val Ala Ile Thr Trp Ala Asp Arg Ala Phe Pro Ser Leu Ala Trp Ile Met Pro Phe Ala Ile Ser Thr Ser Leu Phe Ser Asn Leu Leu Ile Ser Ile Phe Lys Ser Ser Arg Pro Ile Tyr Leu Ala Ser Gln Glu Gly Gln Leu Pro Leu Leu Phe Asn Thr Leu Asn Ser His Ser Ser Pro Phe Thr Ala Val Leu Leu Val Thr Leu Gly Ser Leu Ala Ile Ile Leu Thr Ser Leu Ile Asp Leu Ile Asn Tyr Ile Phe Phe Thr Gly Ser Leu Trp Ser Ile Leu Leu Met Ile Gly Ile Leu Arg Arg Tyr Gln Glu Pro Asn Leu Ser Ile Pro Tyr Lys Val Lys Leu Asp Phe <210> 62 <211> 348 <212> PRT <213> Homo sapiens

Met Ala Ala Thr Leu Gly Pro Leu Gly Ser Trp Gln Gln Trp Arg Arg

<400> 62

1				5					10					15	
Cys	Leu	Ser	Ala	Arg	Asp	Gly	Ser	Arg	Met	Leu	Leu	Leu	Leu	Leu	Leu
			20					25				٠	30	•	
Leu	Gly	Ser	Gly	Gln	Gly	Pro	Gln	Gln	Val	Gly	Ala	Gly	Gln	Thr	Phe
		35					40					45			
Glu	Tyr	Leu	Lys	Arg	Glu	His	Ser	Leu	Ser	Lys	Pro	Tyr	Gln	Gly	Val
	50					55					60				
Gly	Thr	Gly	Ser	Ser	Ser	Leu	Trp	Asn	Leu	Met	Gly	Asn	Ala	Met	Val
65					70					75					80
Met	Thr	G1n	Tyr	Ile	Arg	Leu	Thr	Pro	Asp	Met	Gln	Ser	Lys	Gln	Gly
				85					90					95	
Ala	Leu	Trp	Asn	Arg	Val	Pro	Cys	Phe	Leu	Arg	Asp	Trp	Glu	Leu	Gln
			100					105			•		110		
Val	His	Phe	Lys	Ile	His	Gly	Gln	Gly	Lys	Lys	Asn	Leu	His	Gly	Asp
		115					120					125			
Gly	Leu	Ala	Ile	Trp	Tyr	Thr	Lys	Asp	Arg	Met	Gln	Pro	Gly	Pro	Val
	130					135	•				140				
Phe	Gly	Asn	Met	Asp	Lys	Phe	Val	Gly	Leu	Gly	Val	Phe	Val	Asp	Thr
145	•				150	1	•			155)				160
Tyr	Pro	Asn	Glu	Glu	Lys	Gln	Gln	Glu	Arg	Val	Phe	Pro	Tyr	Ile	Ser
				165	;				170)				175	;
Ala	Met	. Val	. Asn	Asn	Gly	Ser	Leu	Ser	Туз	Asp	His	Glu	. Arg	Asp	Gly
			180)				185	5				190)	
Are	Pro	The	Glu	ı Let	ı G13	Gly	Cys	Thi	r Ala	a Ile	e Val	Arg	g Asr	Leu	Hi
		195	5				200)				205	5		

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[vr /	Asp	Thr	Phe	Leu	Val	Ile	Arg	Tyr	Val	Lys	Arg	His	Leu	Thr	Ile
	210					215					220				
		Asp	Tle	Asp	Glv		His	Glu	Trp	Arg	Asp	Cys	Ile	Glu	Val
	nic c	пор	110		230	-,-				235					240
225	01 .	V-1	A	Leu		Ara	Clv	Tvr	Tvr		Glv	Thr	Ser	Ser	Ile
Pro	GIÀ	Val	WLR		110	vr P	O1,	.,.	250	• • • •	,			255	
				245	A	A	uic	Asn		Ιlο	Ser	l.eu	l.vs		Phe
Thr	Gly	Asp		Ser	Asp	ASI	піѕ			116	Jei	Dog	270		
			260					265		C1		Lou			Aen
Glu	Leu	Thr	Val	Glu	Arg	Thr			Glu	GIU	Lys			VI.R	vsh
		275					280					285			41.
Val	Phe	Leu	Pro	Ser	Val	Asp	Asn	Met	Lys	Leu			Met	Ihr	. VIS
	290	•				299					300				
Pro	Leu	Pro	Pro	Leu	Ser	G1;	y Leu	ı Ala	Let	ı Phe	e Lei	ı Ile	Ya]	Ph€	e Phe
305					310)				319	5				320
Ser	Le	ı Va	l Ph	e Sei	r Val	l Ph	e Ala	a Ile	e Va	l Il	e Gl	y Ile	e Ile	e Lei	ty1
				329	5				33	0				33	5
Asn	Ly	s Tr	p G1	n Gl	u Gl	n Se	r Ar	g Ly	s Ar	g Ph	е Ту	r			
			34	0				34	5						
<2 2	10>	63													
<2	11>	261													
⟨2	12>	PRT													

Met Glu Leu Leu Gln Val Thr Ile Leu Phe Leu Leu Pro Ser Ile Cys

<213≻ Homo sapiens

<400> 63

1				5					10					15	
Ser	Ser	Asn	Ser	Thr	Gly	Val	Leu	Glu	Ala	Ala	Asn	Asn	Ser	Leu	Val
			20	·	,	•	•	25	•				30		
Val	Thr	Thr	Thr	Lys	Pro	Ser	Ile	Thr	Thr	Pro	Asn	Thr	Glu	Ser	Leu
		35					40					45			
Gln	Lys	Asn	Val	Val	Thr	Pro	Thr	Thr	Gly	Thr	Thr	Pro	Lys	Gly	Thr
	50					55		•			60				
Ile	Thr	Asn	Glu	Leu	Leu	Lys	Met	Ser	Leu	Met	Ser	Thr	Ala	Thr	Phe
65					70					75					80
Leu	Thr	Ser	Lys	Asp	Glu	Gly	Leu	Lys	Ala	Thr	Thr	Thr	Asp	Val	Arg
				85					90					95	
Lys	Asn	Asp	Ser	Ile	Ile	Ser	Asn	Val	Thr	Val	Thr	Ser	Val	Thr	Leu
			100	•				105					110)	
Pro	Asn	Ala	Val	Ser	Thr	Leu	Gln	Ser	Ser	Lys	Pro	Lys	Thr	Glu	Thr
		115	,		•		120	ı				125	5		
Gln	Ser	Ser	· Ile	Lys	Thr	Thr	Glu	Ile	Pro	Gly	Ser	· Val	Leu	Glr	Pro
	130)				135	;				140)			
Asp	Ala	Ser	r Pro	Ser	Lys	Thr	· Gly	Thr	Leu	Thi	Ser	: Ile	e Pro	Va]	Thr
145	;				150)				15	5				160
Ile	Pro	Glu	ı Ası	n Thi	Ser	Glr	Ser	Gln	val	. 11	e Gly	y Thi	r Glu	ı Gly	y Gly
				168	5				170)				179	5
Lys	s Asr	n Ala	a Se	r Thi	r Sei	r Ala	a Thi	: Ser	. Ar	g Se	r Ty	r Se	r Se	r Ile	e Ile
			18	0				185	5				19	0	
Leu	ı Pro	o Va	l Va	1 II	e Ala	a Lei	ı Ile	e Val	l Il	e Th	r Le	u Se	r Va	1 Ph	e Va
		19	5				20	0				20	5		

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Leu Val Gly Leu Tyr Arg Met Cys Trp Lys Ala Asp Pro Gly Thr Pro Glu Asn Gly Asn Asp Gln Pro Gln Ser Asp Lys Glu Ser Val Lys Leu Leu Thr Val Lys Thr Ile Ser His Glu Ser Gly Glu His Ser Ala Gln Gly Lys Thr Lys Asn (210) 64 <211> 222 <212> PRT <213> Homo sapiens <400> 64 Met Leu Trp Leu Leu Phe Phe Leu Val Thr Ala Ile His Ala Glu Leu Cys Gln Pro Gly Ala Glu Asn Ala Phe Lys Val Arg Leu Ser Ile Arg Thr Ala Leu Gly Asp Lys Ala Tyr Ala Trp Asp Thr Asn Glu Glu Tyr Leu Phe Lys Ala Met Val Ala Phe Ser Met Arg Lys Val Pro Asn Arg Glu Ala Thr Glu Ile Ser His Val Leu Leu Cys Asn Val Thr Gln Arg Val Ser Phe Trp Phe Val Val Thr Asp Pro Ser Lys Asn His Thr Leu

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Pro Ala Val Glu Val Gln Ser Ala Ile Arg Met Asn Lys Asn Arg Ile Asn Asn Ala Phe Phe Leu Asn Asp Gln Thr Leu Glu Phe Leu Lys Ile Pro Ser Thr Leu Ala Pro Pro Met Asp Pro Ser Val Pro Ile Trp Ile Ile Ile Phe Gly Val Ile Phe Cys Ile Ile Ile Val Ala Ile Ala Leu Leu Ile Leu Ser Gly Ile Trp Gln Arg Arg Arg Lys Asn Lys Glu Pro Ser Glu Val Asp Asp Ala Glu Asp Lys Cys Glu Asn Met Ile Thr Ile Glu Asn Gly Ile Pro Ser Asp Pro Leu Asp Met Lys Gly Gly His Ile Asn Asp Ala Phe Met Thr Glu Asp Glu Arg Leu Thr Pro Leu <210> 65 <211> 183 <212> PRT <213> Homo sapiens <400> 65

Met Gly Val Arg Val His Val Val Ala Ala Ser Ala Leu Leu Tyr Phe

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Ile	Leu	Leu	Ser	Gly	Thr	Arg	Cys	Glu	Glu	Asn	Cys	Gly	Asn	Pro	Glu
			20					25					30		
His	Cys	Leu	Thr	Thr	Asp	Trp	Val	His	Leu	Trp	Tyr	Ile	Trp	Leu	Leu
		35					40					45			
Val	Val	Ile	Gly	Ala	Leu	Leu	Leu	Leu	Cys	Gly	Leu	Thr	Ser	Leu	Cys
	50					55					60				
Phe	Arg	Cys	Cys	Cys	Leu	Ser	Arg	Gln	Gln	Asn	Gly	Glu	Asp	Gly	Gly
65					70					75					80
Pro	Pro	Pro	Cys	Glu	Val	Thr	Val	Ile	Ala	Phe	Asp	His	Asp	Ser	Thr
				85					90					95	
Leu	Gln	Ser	Thr	Ile	Thr	Ser	Leu	Gln	Ser	Val	Phe	Gly	Pro	Ala	Ala
			100					105					110		
Arg	Arg	Ile	Leu	Ala	Val	Ala	His	Ser	His	Ser	Ser	Leu	Gly	Gln	Leu
		115					120					125			
Pro	Ser	Ser	Leu	Asp	Thr	Leu	Pro	Gly	Tyr	Glu	Glu	Ala	Leu	His	Met
	130					135					140				
Ser	Arg	Phe	Thr	Val	Ala	Met	Cys	Gly	Gln	Lys	Ala	Pro	Asp	Leu	Pro
145	•				150	ı				155	,				160
Pro	Val	Pro	Glu	Glu	Lys	Gln	Leu	Pro	Pro	Thr	Glu	Lys	Glu	Ser	Thr
				165	;				170)				175	;
Arg	Ile	Val	. Asp	Ser	Trp	Asn	1								
			180)											

⟨210⟩ 66

<211> 262

<212	> PR	T													
<213> Homo sapiens															
<400	> 66	6												٠	
Met	Gly	Lys	Thr	Phe	Ser	Gln	Leu	Gly	Ser	Trp	Arg	Glu	Asp	Glu	Asn
1				5					10					15	
Lys	Ser	Ile	Leu	Ser	Ser	Lys	Pro	Ala	Ile	Gly	Ser	Lys	Ala	Val	Asn
			20					25					30		
Tyr	Ser	Ser	Thr	Gly	Ser	Ser	Lys	Ser	Phe	Cys	Ser	Cys	Val	Pro	Cys
		35					40					45			
Glu	Gly	Thr	Ala	Asp	Ala	Ser	Phe	Val	Thr	Cys	Pro	Thr	Cys	Gln	Gly
	50					55					60				
Ser	Gly	Lys	Ile	Pro	Gln	Glu	Leu	Glu	Lys	Gln	Leu	Val	Ala	Leu	Ile
65					70		•			7 5					80
Pro	Tyr	Gly	Asp	Gln	Arg	Leu	Lys	Pro	Lys	His	Thr	Lys	Leu	Phe	Val
				85					90					95	
Phe	Leu	Ala	Val	Leu	Ile	Cys	Leu	Val	Thr	Ser	Ser	Phe	Ile	Val	Phe
			100					105					110		
Phe	Leu	Phe	Pro	Arg	Ser	Val	Ile	Val	Gln	Pro	Ala	Gly	Leu	Asn	Ser
		115					120				•	125			
Ser	Thr	Val	Ala	Phe	Asp	Glu	Ala	Asp	Ile	Tyr	Leu	Asn	Ile	Thr	Asn
	130					135					140				
Ile	Leu	Asn	Ile	Ser	Asn	Gly	Asn	Tyr	Tyr	Pro	Ile	Met	Val	Thr	Gln
145					150					155					160
Leu	Thr	Leu	Glu	Val	Leu	His	Leu	Ser	Leu	Val	Val	Gly	Gln	Val	Ser
				165					170					175	

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Asn Asn Leu Leu His Ile Gly Pro Leu Ala Ser Glu Gln Met Phe Tyr Ala Val Ala Thr Lys Ile Arg Asp Glu Asn Thr Tyr Lys Ile Cys Thr Trp Leu Glu Ile Lys Val His His Val Leu Leu His Ile Gln Gly Thr Leu Thr Cys Ser Tyr Leu Ser His Ser Glu Gln Leu Val Phe Gln Ser Tyr Glu Tyr Val Asp Cys Arg Gly Asn Ala Ser Val Pro His Gln Leu Thr Pro His Pro Pro <210> 67 <211> 168 <212> PRT <213> Homo sapiens <400> 67 Met Gly Val Pro Thr Ala Leu Glu Ala Gly Ser Trp Arg Trp Gly Ser Leu Leu Phe Ala Leu Phe Leu Ala Ala Ser Leu Gly Lys Asp Ala Pro Ser Asn Cys Val Val Tyr Pro Ser Ser Ser Gln Glu Ser Glu Asn Ile Thr Ala Ala Ala Leu Ala Thr Gly Ala Cys Ile Val Gly Ile Leu Cys

50			55					60				
Leu Pro Leu	Ile Leu	Leu I	Leu	Val	Tyr	Lys	Gln	Arg	G1n	Ala	Ala	Ser
65		70			•		75					80
Asn Arg Arg	Ala Gln	Glu I	Leu	Val	Arg	Met	Asp	Ser	Asn	Ile	Gln	Gly
	85					90					95	
Ile Glu Asn	Pro Gly	Phe (Glu	Ala	Ser	Pro	Pro	Ala	Gln	Gly	Ile	Pro
	100				105					110		
Glu Ala Lys	Val Arg	His	Pro	Leu	Ser	Tyr	Val	Ala	G1n	Arg	Gln	Pro
115				120					125			
Ser Glu Ser	Gly Arg	His	Leu	Leu	Ser	Glu	Pro	Ser	Thr	Pro	Leu	Ser
130			135					140				
Pro Pro Gly	Pro Gly	Asp '	Val	Phe	Phe	Pro	Ser	Leu	Asp	Pro	Val	Pro
145		150					155					160
Asp Ser Pro	Asn Phe	Glu	Val	Ile								
	168	i										
<210> 68												
<211> 243												
<212> PRT												٠
<213> Homo	sapiens											
<400> 68										•		
Met Ser Ser	Gly Th	r Glu	Leu	Leu	Trp	Pro	Gly	Ala	Ala	Leu	Leu	Val
1	!	5				10)				15	
Leu Leu Gly	Val Al	a Ala	Ser	Leu	Cys	: Val	. Arg	Cys	Ser	Arg	Pro	Gly
	20				25					30		

\la l	.ys	Arg	Ser	Glu	Lys	Ile	Tyr	Gln	Gln	Arg	Ser	Leu	Arg	Glu	Asp
		35					40					45			
Gln (Gln	Ser	Phe	Thr	Gly	Ser	Arg	Thr	Tyr	Ser	Leu	Val	Gly	Gln	Ala
	50					55					60				
Trp	Pro	Gly	Pro	Leu	Ala	Asp	Met	Ala	Pro	Thr	Arg	Lys	Asp	Lys	Leu
65	•				70					75					80
Leu	Gln	Phe	Tyr	Pro	Ser	Leu	Glu	Asp	Pro	Ala	Ser	Ser	Arg	Tyr	Gln
				85					90					95	
Asn	Phe	Ser	Lys	Gly	Ser	Arg	His	Gly	Ser	Glu	Glu	Ala	Tyr	Ile	Asp
			100					105					110		
Pro	Ile	Ala	Met	: Glu	Tyr	Tyr	Asn	Trp	Gly	Arg	, Phe	Ser	Lys	Pro	Pro
		115					120					129			
Glu	Asp	Asp	Ası	Ala	Asn	. Ser	Tyr	Glu	ı Asr	ı Val	L Leu	11e	e Cys	Lys	Gln
	130					135					140				
Lvs			r Gli	u Thi	Gly	, Ala	a Glr	ı Glı	ı Glı	ı Gl	y Ile	e G1	y Gl	y Lei	ı Cys
145					150					15					160
	G1,	v As	ъ Le	u Sei	r Lei	u Sei	r Lei	u Ala	a Le	u Ly	s Th	r Gl	y Pr	o Th	r Ser
		,	-	16					17					17	
C1 v	م آ	n Cv	s Pr			a Se	r Pr	o Gl	u Gl	u As	p Gl	u Gl	u Se	r Gl	u Asp
01,			18					18					19		
Tur	· 61	n As			a Se	r Il	e Hi	s Gl	n Tr	p Ar	g Gl	u Se	er Ar	g Ly	s Val
1,71	. 01	19			•		20					20			
Mad	. <u>C</u> 1			au (C1	n Ar	₋ و (۲۵			er Pi	ro Gi	ly Pi	o Va	al Gl	ly Se	er Pr
gle)			rii D	Ju 01		21						20			
	21 - C1		1,, A	en G1	lv G1			sp Ts	yr V:	al A			lu ·Va	al A	la Al

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Thr Glu Ala <210> 69 <211> 428 <212> PRT <213> Homo sapiens <400> 69 Met Ala Arg Ser Leu Cys Pro Gly Ala Trp Leu Arg Lys Pro Tyr Tyr Leu Gln Ala Arg Phe Ser Tyr Val Arg Met Lys Tyr Leu Phe Phe Ser Trp Leu Val Val Phe Val Gly Ser Trp Ile Ile Tyr Val Gln Tyr Ser Thr Tyr Thr Glu Leu Cys Arg Gly Lys Asp Cys Lys Lys Ile Ile Cys Asp Lys Tyr Lys Thr Gly Val Ile Asp Gly Pro Ala Cys Asn Ser Leu Cys Val Thr Glu Thr Leu Tyr Phe Gly Lys Cys Leu Ser Thr Lys Pro Asn Asn Gln Met Tyr Leu Gly Ile Trp Asp Asn Leu Pro Gly Val Val Lys Cys Gln Met Glu Gln Ala Leu His Leu Asp Phe Gly Thr Glu Leu

Glu	Pro	Arg	Lys	Glu	Ile	Val	Leu	Phe	Asp	Lys	Pro	Thr	Arg	Gly	Thr
	130					135					140				
Thr	Val	Gln	Lys	Phe	Lys	Glu	Met	Val	Tyr	Ser	Leu	Phe	Lys	Ala	Lys
145					150					155					160
Leu	Gly	Asp	Gln	Gly	Asn	Leu	Ser	Glu	Leu	Val	Asn	Leu	Ile	Leu	Thr
				165					170					175	
Val	Ala	Asp	Gly	Asp	Lys	Asp	Gly	Gln	Val	Ser	Leu	Gly	Glu	Ala	Lys
			180					185				-	190		
Ser	Ala	Trp	Ala	Leu	Leu	Gln	Leu	Asn	Glu	Phe	Leu	Leu	Met	Val	Ile
		195					200					205			
Leu	Gln	Asp	Lys	Glu	His	Thr	Pro	Lys	Leu	Met	Gly	Phe	Cys	Gly	Asp
	210		٠			215					220				
Leu	Tyr	Val	Met	Glu	Ser	Val	Glu	Tyr	Thr	Ser	Leu	Tyr	Gly	Ile	Ser
225	i				230					235					240
Leu	Pro	Trp	Val	Ile	Glu	Leu	Phe	Ile	Pro	Ser	Gly	Phe	Arg	Arg	Ser
				245					250	١				255	
Met	Asp	Gln	Leu	Phe	Thr	Pro	Ser	Trp	Pro	Arg	Lys	Ala	Lys	Ile	Ala
			260)				265	,				270		
Ile	Gly	/ Leu	ı Leu	Glu	Phe	· Val	Glu	Asp	Val	Phe	His	Gly	Pro	Tyr	Gly
		275	5				280)				285	,		
Ası	n Phe	e Leu	ı Met	Cys	Asp	Thr	Ser	Ala	Lys	Asn	Leu	Gly	Tyr	Asn	Asp
	290)				295	5				300)			
Lys	s Ty	r Ası	Lev	ı Lys	s Met	: Va	l Asp	Met	: Arg	g Lys	Ile	. Val	Pro	Glu	Thr
30	5				310)				315	5				320
Ası	n Lei	u Ly:	s Glu	Leı د	ı Ile	E Ly:	s Asp	Ar _i	g His	s Cys	s Glu	ı Sei	r Ası	Lei	ı Asp

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Cys Val Tyr Gly Thr Asp Cys Arg Thr Ser Cys Asp Gln Ser Thr Met Lys Cys Thr Ser Glu Val Ile Gln Pro Asn Leu Ala Lys Ala Cys Gln Leu Leu Lys Asp Tyr Leu Leu Arg Gly Ala Pro Ser Glu Ile Arg Glu Glu Leu Glu Lys Gln Leu Tyr Ser Cys Ile Ala Leu Lys Val Thr Ala Asn Gln Met Glu Met Glu His Ser Leu Ile Leu Asn Asn Leu Lys Thr Leu Leu Trp Lys Lys Ile Ser Tyr Thr Asn Asp Ser <210> 70 <211> 283 <212> PRT <213> Homo sapiens <400> 70 Met Pro His Ser Ser Leu His Pro Ser Ile Pro Cys Pro Arg Gly His Gly Ala Gln Lys Ala Ala Leu Val Leu Leu Ser Ala Cys Leu Val Thr Leu Trp Gly Leu Gly Glu Pro Pro Glu His Thr Leu Arg Tyr Leu Val

Leu	His	Leu	Ala	Ser	Leu	Gln	Leu	Gly	Leu	Leu	Leu	Asn	Gly	Val	Cys
	50					55					60				
Ser	Leu	Ala	Glu	Glu	Leu	His	His	Ile	His	Ser	Arg	Tyr	Arg	Gly	Ser
65					70					75					80
Tyr	Trp	Arg	Thr	Val	Arg	Ala	Cys	Leu	Gly	Cys	Pro	Leu	Arg	Arg	Gly
				85					90					95	
Ala	Leu	Leu	Leu	Leu	Ser	Ile	Tyr	Phe	Tyr	Tyr	Ser	Leu	Pro	Asn	Ala
			100					105					110		
Val	Gly	Pro	Pro	Phe	Thr	Trp	Met	Leu	Ala	Leu	Leu	Gly	Leu	Ser	Gln
		115					120					125			
Ala	Leu	Asn	Ile	Leu	Leu	Gly	Leu	Lys	Gly	Leu	Ala	Pro	Ala	Glu	Ile
	130					135					140				
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Trp	Ser	Tyr	Tyr	Ile	Gly	Tyr	Leu	Arg	Leu	Ile	Leu	Pro	Glu	Leu	G1n
				165					170					175	
Ala	Arg	Ile	Arg	Thr	Tyr	Asn	Gln	His	Tyr	Asn	Asn	Leu	Leu	Arg	Gly
			180					185					190		
Ala	Val	Ser	Gln	Arg	Leu	Tyr	Ile	Leu	Leu	Pro	Leu	Asp	Cys	Gly	Val
		195					200					205			
Pro	Asp	Asn	Leu	Ser	Met	Ala	Asp	Pro	Asn	Ile	Arg	Phe	Leu	Asp	Lys
	210					215					220				
Leu	Pro	Gln	Gln	Thr	Ala	Asp	Arg	Ala	Gly	Ile	Lys	Asp	Arg	Val	Tyr
225					230					235					240
Ser	Asn	Ser	Ile	Tvr	Glu	Leu	Len	Glu	Asn	Glv	Gln	Aro	Asn	Leu	Gln

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245 250 255

Met Thr Ala Ala Ser Arg Cys Pro Arg Arg Phe Ser Gly Thr Cys Gly

260 265 270

Arg Arg Lys Arg Leu Leu Trp Ala Ala

275 280

<210> 71

⟨211⟩ 1167

<212> DNA

<213> Homo sapiens

<400> 71

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cttaatagtc	actcttctcc	atttacagct	gtgctactac	ttgtcacttt	gggatccctt	1020
gcaattatct	taacaagtct	aattgatttg	ataaactata	ttttttcac	gggttcatta	1080
tggtctatat	tattaatgat	aggaatacta	aggcggagat	accaggaacc	caatctatct	1140
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<210> 72

<211> 1044

<212> DNA

<213> Homo sapiens

<400> 72

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tcagataatc	atgatgtcat	ttccttgaag	ttgtttgaac	tgacagtgga	gagaacccca	840
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gagatgacag	ctccactgcc	gccctgagt	ggcctggccc	tcttcctcat	cgtcttttc	960
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<210> 73

(211) 783

<212> DNA

<213> Homo sapiens

<400> 73

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(210)	14
<211>	666

<212> DNA

<213> Homo sapiens

<400> 74

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<210> 75

<211> 549

<212> DNA

(213) Homo sapiens

<400> 75

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60

158/307

120 gggacgagat gtgaggaaaa ctgtggtaat cctgaacatt gcctgaccac agactgggta 180 catctctggt atatatggtt gctagtggta attggcgcgc tgcttctcct gtgtggcctg 240 acgtccctgt gcttccgctg ctgctgtctg agccgccagc aaaatgggga agatgggggc 300 ccaccacct gtgaagtgac cgtcattgct ttcgatcacg acagcactct ccagagcact 360 atcacatctc tgcagtcggt gtttggccct gcagctcgga ggatcctggc tgtggctcac 420 tcccacaget ccctgggcca gctgccctcc tctttggaca ccctcccagg gtatgaagaa 480 gctcttcaca tgagtcgctt cacagtagcc atgtgcgggc agaaagcacc tgatctaccc ccagtacctg aagaaaagca gctgcctcca acagagaagg agtcgactcg aatagttgac 540 549 tcttggaac

<210> 76

<211> 786

<212> DNA

<213> Homo sapiens

<400> 76

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gatgaaaaca	catacaaaat	ctgtacctgg	ctggaaatca	aagtccacca	tgtgcttttg	660
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agctatgaat	atgtggactg	ccgaggaaac	gcatctgtgc	cccaccagct	gacccctcac	780
ccacca						786

<210> 77

<211> 504

<212> DNA

<213> Homo sapiens

<400> 77

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<210> 78

<211> 729

<212> DNA

<213> Homo sapiens

<400> 78

160/307

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cagcagagaa	gtctgcgtga	ggaccaacag	agctttacgg	ggtcccggaċ	ctactccttg	180
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cagtggcgcg	agtccaggaa	ggtcatgggg	caactccaga	gagaagcatc	ccctggcccg	660
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acagaagcc						729

<210> 79

<211> 1284

<212> DNA

<213> Homo sapiens

<400> 79

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tggattatat atgtgcagta ttctacctat acagaattat gcagaggaaa ggactgtaag 180
aaaataatat gtgacaagta caagactgga gttattgatg ggcctgcatg taacagcctt 240
tgtgttacag aaactcttta ctttggaaaa tgtttatcca ccaagcccaa caatcagatg 300
tatttaggga tttgggataa tctaccaggt gttgtgaaat gtcaaatgga acaagcgctt 360

161/307

a 420	tgataagcca	tagtgctatt	agaaaagaaa	attggaacca	ttggaactga	catcttgatt
ia 480	taaggcaaaa	atagtctctt	gaaatggtct	aaaatttaaa	ctactgtaca	actagaggaa
ga 540	ggctgatgga	tcttgacggt	gttaatctca	ctctgaactg	aaggaaacct	ttgggtgacc
g 600	tcttcaactg	catgggcact	gcaaagtcgg	cttgggagaa	gccaggtttc	gacaaagatg
ga 660	attaatggga	ataccccaa	gataaagaac	gatacttcaa	ttctcatggt	aatgaatttc
gc 720	tggaataagc	cctctcttta	gttgaatata	gatggaaagt	acctctatgt	ttctgtggtg
g 780	ggatcagctg	gaagaagcat	tctgggttca	ttttattcca	tcattgaact	cttccttggg
a 840	atttgtggaa	gacttctaga	atagccatag	aaaggccaaa	catggccaag	ttcacaccat
a 900	caaaaaccta	atactagtgc	ctcatgtgcg	cggaaatttc	atggccccta	gatgttttcc
a 960	gccagagaca	gaaaaattgt	gtggatatga	tttgaaaatg	ataagtatga	ggatataatg
gc 1020	tgtctatggc	atttggactg	tgtgagtctg	ggatcgtcac	aacttattaa	aacctgaaag
ia 1080	agtgatacaa	gtacttcaga	acaatgaagt	tgatcagagt	gaactagctg	acagattgta
t 1140	tgctccaagt	tactgcgtgg	aaagactacc	tcagttactc	caaaagcttg	ccaaacttgg
a 1200	agtcacagca	ttgctctcaa	tattcttgta	aaagcagctt	aagaattaga	gaaattcgtg
ig 1260	attgtggaag	taaaaacatt	ctaaataacc	ttctttgata	aaatggaaca	aatcaaatgg
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<210> 80

<211> 849

<212> DNA

<213> Homo sapiens

<400> 80

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ctgtccatct	atttctacta	ctecctccca	aatgcggtcg	gcccgccctt	cacttggatg	360
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ccagctgaga	tctctgcagt	gtgtgaaaaa	gggaatttca	acgtggccca	tgggctggca	480
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⟨210⟩ 81

⟨211⟩ 1376

<212> DNA

<213≻ Homo sapiens

<220>

. <221> CDS

⟨222⟩ (100)...(1269)

<400> 81

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Met Asp Arg Gly Glu

1

888	ata	cag	ctc	aag	aga	gtg	ttt	gga	tat	tgg	tgg	ggc	aca	agt	ttt	162
Lys	Ile	Gln	Leu	Lys	Arg	Val	Phe	Gly	Tyr	Trp	Trp	Gly	Thr	Ser	Phe	
				10					15					20		
ttg	ctt	att	aat	atc	att	ggt	gca	gga	att	ttt	gtg	tcc	ccc	aaa	ggt	210
Leu	Leu	Ile	Asn	Ile	Ile	Gly	Ala	Gly	Ile	Phe	Val	Ser	Pro	Lys	Gly	
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gtg	ttg	gca	tac	tct	tgc	atg	aac	gtg	gga	gtc	tcc	ctg	tgc	gtt	tgg	258
Val	Leu	Ala	Tyr	Ser	Cys	Met	Asn	Val	Gly	Val	Ser	Leu	Cys	Val	Trp	
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gct	ggc	tgt	gcc	ata	ctg	gcc	atg	aca	tca	act	ctt	tgc	tct	gca	gag	306
Ala	Gly	Cys	Ala	Ile	Leu	Ala	Met	Thr	Ser	Thr	Leu	Cys	Ser	Ala	Glu	
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Arg	Tyr	Phe	Gly	Ser	Thr	Val	Ala	Phe	Leu	Asn	Leu	Trp	Thr	Ser	Leu	
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Phe	Leu	G1y	Ser	Gly	Val	Val	Ala	Gly	Gln	Ala	Leu	Leu	Leu	Ala	Glu	
			105					110	•				115			
tac	agc	atc	cag	cct	ttt	ttt	ccc	agc	tgc	tct	gtc	сса	aag	ctg	cct	498
Tyr	Ser	Ile	Gln	Pro	Phe	Phe	Pro	Ser	Cys	Ser	Val	Pro	Lys	Leu	Pro	
		120					125					130				
888	aaa	tet	ctg	gca	ttø	gcc	atg	tte	tgg	att	ota	oos	att	ctg	act	546

Lys	Lys	Cys	Leu	Ala	Leu	Ala	Met	Leu	Trp	Ile	Val	Gly	Ile	Leu	Thr	
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Ser	Arg	Gly	Val	Lys	Glu	Val	Thr	Trp	Leu	Gln	Ile	Ala	Ser	Ser	Val	
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Leu	Ile	Arg	Gly	Lys	Lys	Glu	Asn	Val	Glu	Arg	Phe	Gln	Asn	Ala	Phe	
			185					190					195			
gat	gct	gaa	ctt	cca	gat	atc	tct	cac	ctt	ata	caa	gcc	atc	ttc	caa	738
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ccc	aaa	tgc	ata	ttt	act	gcg	tta	cct	ctg	gtg	act	gta	gtt	tat	tta	834
Pro	Lys	Cys	Ile	Phe	Thr	Ala	Leu	Pro	Leu	Val	Thr	Val	Val	Tyr	Leu	
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Leu	Val	Asn	Ile	Ser	Tyr	Leu	Thr	Val	Leu	Thr	Pro	Arg	Glu	Ile	Leu	
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tct	tca	gat	gct	gta	gct	atc	aca	tgg	gct	gat	cga	gct	ttt	ccc	tca	930
Sar	Sar	Acn	Δla	Va 1	Δla	Tla	Thr	Trn	Ala	Acn	Ara	Δla	Pho	Pro	Ser	

			265					270					275			
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Leu	Ala	Trp	Ile	Met	Pro	Phe	Ala	Ile	Ser-	Thr	Ser	Leu	Phe	Ser	Asn	
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Tyr	Ģln	Glu	Pro	Asn	Leu	Ser	Ile	Pro	Tyr	Lys	Val	Lys	Leu	Asp	Phe	
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Phe Tyr

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Thr	Asn	Glu	Leu	Leu	Lys	Met	Ser	Leu	Met	Ser	Thr	Ala	Thr	Phe	Leu	
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					Ala										_	
		180					185	6		-,-		190				
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					Leu											
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As	р Ме	t Ly	s Gl	y Gl	y Hi	s Il	e As	n As	p Al	a Ph	e Me	t Th	ır Gl	u As	p Gl	u	
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Ar	g Le	eu Th	nr Pi	ro Le	u												
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Leu Leu Tyr Phe Ile Leu Leu Ser Gly Thr Arg Cys Glu Glu Asn Cys

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Ile	Trp	Leu	Leu	Val	Val	Ile	Gly	Ala	Leu	Leu	Leu	Leu	Cys	Gly	Le	eu		
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acg	tcc	ctg	tgc	ttc	cgc	tgc	tgc	tgt	ctg	agc	cgc	cag	caa	aat	g	gg	30)2
Thr	Ser	Leu	Cys	Phe	Arg	Cys	Cys	Cys	Leu	Ser	Arg	Gln	Gln	Asn	ı G	ly		
				65					70					75	5			
gaa	gat	ggg	ggc	cca	cca	ccc	tgt	gaa	gtg	acc	gto	att	gct	tto	g	at	38	50
Glu	ı Asp	Gly	Gly	Pro	Pro	Pro	Cys	Glu	Val	Thr	Val	Ile	e Ala	a Phe	e A	sp		
			80)				85					90)				
				cto													. 3	98
Hi	s Ası	Se:	r Thi	r Leu	Glr	Ser	Thr	Ile	Thr	Ser	Let	ı Gl	n Se	r Va	1 F	he		
		9	5				100)				10	5					
gg	c cc	t gc	a gc	t cgg	g agg	g atc	ctg	g gct	t gtg	g gct	ca.	c tc	c ca	c ag	(C 1	tcc	4	46
G1	y Pr	o Al	a Al	a Ar	g Arı	g Ile	e Lei	ı Ala	a Val	l Ala	a Hi	s Se	r Hi	s Se	r	Ser		
	11	0				115	5				12	0						
				g cc													4	194
Le	u Gl	y Gl	n Le	u Pr	o Se	r Sei	r Le	u As	p Th	r Le	u Pr	o G1	у Ту	r G	lu			
12	25				13	0				13	5					140		
				g ag													;	542
A.	la Le	eu Hi	is Me	et Se	er Ar	g Ph	e Th	r Va	l Al	a Me	t Cy	ys G	ly G			Ala		
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cct gat cta ccc cca gta cct gaa gaa aag cag ctg cct cca aca gag	590
Pro Asp Leu Pro Pro Val Pro Glu Glu Lys Gln Leu Pro Pro Thr Glu	
160 165 170	
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Lys Glu Ser Thr Arg Ile Val Asp Ser Trp Asn	
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<220>

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<222> (236)...(1024)

<400> 86

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tca	atc	ctg	tcc	tcc	aaa	cca	gcc	att	ggc	agc	aag	gct	gtc	aac	tac	334
Ser	Ile	Leu	Ser	Ser	Lys	Pro	Ala	Ile	Gly	Ser	Lys	Ala	Val	Asn	Tyr	
		20					25					30				
tcc	agc	acc	ggt	agc	agc	aag	tct	ttt	tgt	tcc	tgt	gtg	cct	tgt	gaa	382
Ser	Ser	Thr	Gly	Ser	Ser	Lys	Ser	Phe	Cys	Ser	Cys	Val	Pro	Cys	Glu	
	35					40					45					
gga	act	gct	gat	gcc	agc	ttc	gtg	act	tgt	ccc	acc	tgc	cag	ggc	agt	430
Gly	Thr	Ala	Asp	Ala	Ser	Phe	Val	Thr	Cys	Pro	Thr	Cys	G1n	Gly	Ser	
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ggc	aag	att	ccc	caa	gag	ctg	gag	aag	cag	ttg	gtg	gct	ctc	att	ccc	478
Gly	Lys	Ile	Pro	Gln	Glu	Leu	Glu	Lys	Gln	Leu	Val	Ala	Leu	Ile	Pro	
				70					75					80		
tat	ggg	gac	cag	agg	ctg	aag	ссс	aag	cac	acg	aag	ctc	ttt	gtg	ttc	526
Tyr	Gly	Asp	Gln	Arg	Leu	Lys	Pro	Lys	His	Thr	Lys	Leu	Phe	Val	Phe	
			85			٠		90					95			
ctg	gcc	gtg	ctc	ato	tgc	ctg	gtg	acc	tcc	tcc	ttc	ato	gtc	ttt	ttc	574
Leu	Ala	Val	Leu	Ile	Cys	Leu	Val	Thr	Ser	Ser	Phe	Ile	. Val	Phe	Phe	
		100	ı				105					110)			
ctg	ttt	ccc	cgg	tcc	gto	att	gtg	cag	cct	gca	ggo	cto	aac	tco	tcc	622
Leu	Phe	Pro	Arg	Ser	· Val	Ile	Val	Gln	Pro	Ala	Gly	, Lei	ı Asr	Ser	Ser	
	115	;				120)				125	5				
aca	gtg	gcc	ttt	gat	t gag	g gct	gat	ato	tac	cto	820	ata	a ac	g aa1	t atc	670
Thr	· Val	Als	. Phe	. Ası	o Glu	ı Ala	Ast	Ile	. Tvi	Leu	Ası	n Ile	e Thi	. Ası	n Ile	

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Asn	Ile	Ser	Asn	Gly	Asn	Tyr	Tyr	Pro	Ile	Met	Val	Thr	Gln	Leu	
			150					155					160		
ctc	gag	gtt	ctg	cac	ctg	tcc	ctc	gtg	gtg	ggg	cag	gtt	tcc	aac	766
Leu	Glu	Val	Leu	His	Leu	Ser	Leu	Val	Val	Gly	Gln	Val	Ser	Asn	
		165					170					175			
ctt	ctc	cta	cac	att	ggc	cct	ttg	gcc	agt	gaa	cag	atg	ttt	tac	814
Leu	Leu	Leu	His	Ile	Gly	Pro	Leu	Ala	Ser	Glu	Gln	Met	Phe	Tyr	
	180					185					190				
gta	gct	acc	aag	ata	cgg	gat	gaa	aac	aca	tac	aaa	atc	tgt	acc	862
Val	Ala	Thr	Lys	Ile	Arg	Asp	Glu	Asn	Thr	Tyr	Lys	Ile	Cys	Thr	
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ctg	gaa	atc	aaa	gtc	cac	cat	gtg	ctt	ttg	cac	ato	cag	ggo	acc	910
Leu	Glu	Ile	Lys	Val	His	His	Val	Leu	Leu	His	Ile	Glr	Gly	Thr	
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acc	tgt	tca:	tac	ctg	ago	cat	. tca	gag	cag	ctg	gto	tt1	t cas	g agc	958
Thr	Cys	Ser	Tyr	Leu	Ser	His	Ser	Glu	Glr	Leu	ı Val	Phe	e Gla	n Ser	
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· Glu	ı Tyı	r Val	l Ası	Cys	. Ar	g G1;	, Ası	n Ala	a Ser	r Va	l Pro	o Hi	s Gl	n Leu	
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cc.	t ca	c cca	a cca	a tga	acct	gtc	tgcti	gtcc	ct g	tact	ccag	g ca	cctg	caac	106
. Pr	o Hi	s Pro	o Pro	0											
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260

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⟨222⟩ (103)... (609)

<400> 87

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Met Gly Val Pro

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Thr	Ala	Leu	Glu	Ala	Gly	Ser	Trp	Arg	Trp	Gly	Ser	Leu	Leu	Phe	Ala	
5					10			•		15					20	
ctc	ttc	ctg	gct	gcg	tcc	cta	ggc	aaa	gat	gca	cca	tcc	aac	tgt	gtg	210
Leu	Phe	Leu	Ala	Ala	Ser	Leu	Gly	Lys	Asp	Ala	Pro	Ser	Asn	Cys	Val	
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gtg	tac	cca	tcc	tcc	tcc	cag	gag	agt	gaa	aac	atc	acg	gct	gca	gcc	258
Val	Tyr	Pro	Ser	Ser	Ser	Gln	Glu	Ser	Glu	Asn	Ile	Thr	Ala	Ala	Ala	
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ctg	gct	acg	ggt	gcc	tgc	atc	gta	gga	atc	ctc	tgc	ctc	ccc	ctc	atc	306
Leu	Ala	Thr	Gly	Ala	Cys	Ile	Val	Gly	Ile	Leu	Cys	Leu	Pro	Leu	Ile	
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ctg	ctc	ctg	gtc	tac	aag	caa	agg	cag	gca	gcc	tcc	aac	cgc	cgt	gcc	354
Leu	Leu	Leu	Val	Tyr	Lys	Gln	Arg	Gln	Ala	Ala	Ser	Asn	Arg	Arg	Ala	
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cag	gag	ctg	gtg	cgg	atg	gac	agc	aac	att	caa	ggg	att	gaa	aac	ccc	402
Gln	Glu	Leu	Val	Arg	Met	Asp	Ser	Asn	Ile	Gln	Gly	Ile	Glu	Asn	Pro	
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agg	cac	ccc	ctg	tcc	tat	gtg	gco	cag	cgg	cag	cct	tc1	t gag	g tct	ggg	498
Arg	His	Pro	Leu	Ser	Туз	· Val	Ala	Gln	Arg	Glr	Pro	Se	r Glu	ı Sei	Gly	
			120)				125	5				130)		
000		cto	z ctt	tes	7 090	7 000	. 200	900		cts	z toi	t cc	t cc	a ggo	ccc	546

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tgagattctc ccctagagac ctgaaattca ccagctacag atgccaaatg acttacatct	890
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atgtggcagc atcagtggga caagatggac actgggccac cctcccaggc accagacaca	1010
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gggtctgttc ctagttgcaa cagttcttgg aaacccactc gagagggcca cgcctccatt	180
caccaggcca cgcatcacaa gaggcaacac caggagccaa c atg agc tcg ggg	233 .
Met Ser Ser Gly	
. 1	
act gaa ctg ctg tgg ccc gga gca gcg ctg ctg gtg ctg ttg ggg gtg	281
Thr Glu Leu Leu Trp Pro Gly Ala Ala Leu Leu Val Leu Leu Gly Val	
5 10 15 20	
gca gcc agt ctg tgt gtg cgc tgc tca cgc cca ggt gca aag agg tca	329
Ala Ala Ser Leu Cys Val Arg Cys Ser Arg Pro Gly Ala Lys Arg Ser	
25 30 35	
gag aaa atc tac cag cag aga agt ctg cgt gag gac caa cag agc ttt	377
Glu Lys Ile Tyr Gln Gln Arg Ser Leu Arg Glu Asp Gln Gln Ser Phe	
40 45 50	
acg ggg tcc cgg acc tac tcc ttg gtc ggg cag gca tgg cca gga ccc	425
Thr Gly Ser Arg Thr Tyr Ser Leu Val Gly Gln Ala Trp Pro Gly Pro	

		55					60					65				
ctg	gcg	gac	atg	gca	ccc	aca.	agg	aag	gac	aag	ctg	ttg	caa	ttc	tac	473
Leu	Ala	Asp	Met	Ala	Pro	Thr	Arg	Lys	Asp	Lys	Leu	Leu	Gln	Phe	Tyr	
	70					75					80					
ccc	agc	ctg	gag	gat	cca	gca	tct	tcc	agg	tac	cag	aac	ttc	agc	aaa	521
Pro	Ser	Leu	Glu	Asp	Pro	Ala	Ser	Ser	Arg	Tyr	Gln	Asn	Phe	Ser	Lys	
85					90					95					100	
gga	agc	aga	cac	ggg	tcg	gag	gaa	gcc	tac	ata	gac	ccc	att	gcc	atg	569
Gly	Ser	Arg	His	Gly	Ser	Glu	Glu	Ala	Tyr	Ile	Asp	Pro	Ile	Ala	Met	
				105					110					115		
gag	tat	tac	aac	tgg	ggg	cgg	ttc	tcg	aag	ccc	сса	gaa	gat	gat	gat	617
Glu	Tyr	Tyr	Asn	Trp	Gly	Arg	Phe	Ser	Lys	Pro	Pro	Glu	Asp	Asp	Asp	
			120	i				125	i				130)		
gcc	aat	tco	tac	gag	aat	gtg	cto	att	tgc	aag	cag	g aaa	acc	aca	gag	665
Ala	Asn	Sei	. Tyr	Glu	ı Asn	Val	Leu	Ile	. Cys	Lys	Glr	ı Lys	Thr	Thr	Glu	
		13	5				140)		,		145	5			
aca	ggt	t gc	c cas	g cag	g gag	ggg	ata	ggt	ggo	cto	tge	c aga	a ggg	g gao	ctc	713
Thi	Gly	y Ala	a Glı	ı Glı	n Glu	ı Gly	/ Ile	e Gly	/ Gly	y Let	ı Cy:	s Ar	g Gly	y Asj	Leu	
	150	0				158	5				16	0				
age	c ct	g tc	a ct	g gc	c ct	g aag	g act	t gg	c cc	c ac	t tc	t gg	t ct	c tg	t ccc	761
Se	r Le	u Se	r Le	u Al	a Lei	u Ly:	s Thi	r Gl	y Pr	o Th	r Se	r Gl	y Le	u Cy	s Pro	
16	5				17	0				17	5				180	
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Se	r Al	a Se	r Pr	o Gl	u Gl	u As	p Gl	u Gl	u Se	r Gl	u As	р Ту	r Gl		n Ser	
				18	5				19	0				19	5	

gca tcc atc cat cag tgg cgc gag tcc agg aag gtc atg ggg caa ctc	857
Ala Ser Ile His Gln Trp Arg Glu Ser Arg Lys Val Met Gly Gln Leu	
200 205 210	•
cag aga gaa gca tcc cct ggc ccg gtg gga agc cca gac gag gag gac	905
Gln Arg Glu Ala Ser Pro Gly Pro Val Gly Ser Pro Asp Glu Glu Asp	ı
215 220 225	
ggg gaa ccg gat tac gtg aat ggg gag gtg gca gcc aca gaa gcc	950
Gly Glu Pro Asp Tyr Val Asn Gly Glu Val Ala Ala Thr Glu Ala	
230 235 240	
tagggcagac caagaagaaa ggagccaagg caaagaggga ccactgtgct catggac	cca 1010
tcgctgcctt ccaaggacca tttcccagag ctactcaact tttaagcccc tgccatgg	gtt 1070
gctcctggaa ggagaaccag ccaccctgag gaccacctgg ccatgcgtgc acagcctg	ggg 1130.
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gatttaggat aagetgteac ceagteecea taacaaaace aetgteeaac aetggta	tct 1310
gtgttctttt gtgctatgaa tttggattcc taattgctat tgttggttgc tggggtt	tta 1370
aatgattgat aagcttgtac agttaactta tagaggggga gccatattta acattct	gga 1430
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getteagtte acteaggaag aaatgagget gtegeeatet ttatgtgett eeagtgg	aaa 1610
tgtcacttgc tacagacaat agtgcatgag agtctagaga agtagtgacc agaacag	ggc 1670
agagtaggtc ccctccatgg ccctgaatcc tcctctgctc cagggctggc ctctgca	gag 1730
ctgattaaac agtgttgtga ctgtctcatg ggaagagctg gggcccagag ggacctt	gag 1790
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agttg	1855

⟨210⟩ 89	
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Met Ala Arg Ser Leu Cys Pro Gly	
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gcc tgg cta agg aaa ccc tat tac ctc cag gct cgc ttc tca tat gtg	99
Ala Trp Leu Arg Lys Pro Tyr Tyr Leu Gln Ala Arg Phe Ser Tyr Val	
10 15 20	
cgg atg aaa tat ctt ttc ttt tcc tgg tta gtg gtt ttt gtt gga agc	147
Arg Met Lys Tyr Leu Phe Phe Ser Trp Leu Val Val Phe Val Gly Ser	
25 30 35 40	
tgg att ata tat gtg cag tat tct acc tat aca gaa tta tgc aga gga	195
Trp Ile Ile Tyr Val Gln Tyr Ser Thr Tyr Thr Glu Leu Cys Arg Gly	
45 50 55	
aag gac tgt aag aaa ata ata tgt gac aag tac aag act gga gtt att	243
Lys Asp Cys Lys Lys Ile Ile Cys Asp Lys Tyr Lys Thr Gly Val Ile	
60 65 70	
gat ggg cct gca tgt aac agc ctt tgt gtt aca gaa act ctt tac ttt	291

Asp	Gly	Pro	Ala	Cys	Asn	Ser	Leu	Cys	Val	Thr	Glu	Thr	Leu	Tyr	Phe	
		75					80					85		•		
gga	aaa	tgt	tta	tcc	acc	aag	ccc	aac	aat	cag	atg	tat	tta	ggg	att	339
Gly	Lys	Cys	Leu	Ser	Thr	Lys	Pro	Asn	Asn	Gln	Met	Tyr	Leu	Gly	Ile	
	90					95					100					
tgg	gat	aat	cta	cca	ggt	gtt	gtg	aaa	tgt	caa	atg	gaa	caa	gcg	ctt	387
				Pro												
105					110					115					120	
		gat	ttt	gga	act	gaa	ttg	gaa	cca	aga	aaa	gaa	ata	gtg	cta	435
				Gly												
		•		125					130					135		
tt1	gat	t aas	z cca	a act	aga:	gga	act	act	gta:	caa	aaa	ttt	aaa	gaa	atg	483
			•												ı Met	
		, ,,	140			, ,		145					150			,
atı	r ta	t ao			t aas	g gca	a aaa	a tt:	g ggt	t gad	caa	a ġga	a aa	c cto	tct	531
															ı Ser	
٧a	1 17	15		.	· -,		160		•			169				
	+			t ot	c at	c tt			a ac.	t ga	t gg	a ga	c aa	a ga	t ggo	579
															p Gly	
GI			ii ns	II LE	u II	17					18				-	
	17							a to	e ac	a to			t ct	t ca	a cti	g 627
															a cta n Lea	
		al Se	er Le	eu Gl			.a Ly	'S 36	SL WI			a Le	u D	,u	n Le: 20	
	35				19					19				nt 01		
															C CC	
A	sn G	lu P	he L	eu Lo	eu Me	et Va	al I	le L	eu G	In As	sp L	ys G	lu H	15 I	hr Pr	U

205		210	215	
aaa tta atg gga ttc	tgt ggt gac ctc	tat gtg atg gaa	agt gtt gaa 7	723
Lys Leu Met Gly Phe	Cys Gly Asp Leu	Tyr Val Met Glu	Ser Val Glu	
220	225		230	
tat acc tct ctt tat	gga ata agc ctt	cct tgg gtc att	gaa ctt ttt	771
Tyr Thr Ser Leu Tyr	Gly Ile Ser Leu	Pro Trp Val Ile	Glu Leu Phe	
235	240	245		
att cca tct ggg ttc	aga aga agc at	g gat cag ctg ttc	aca cca tca	819
Ile Pro Ser Gly Phe	Arg Arg Ser Me	t Asp Gln Leu Phe	Thr Pro Ser	
250	255	260		
tgg cca aga aag gcc	aaa ata gcc at	a gga ctt cta gaa	ttt gtg gaa	867
Trp Pro Arg Lys Ala	Lys Ile Ala Il	e Gly Leu Leu Glu	Phe Val Glu	
265	270	275	280	
gat gtt ttc cat ggc	ccc tac gga aa	t ttc ctc atg tg	e gat act agt	915
Asp Val Phe His Gly	Pro Tyr Gly As	on Phe Leu Met Cy	s Asp Thr Ser	
285	;	290	295	
gcc aaa aac cta gga	tat aat gat aa	ag tat gat ttg aa	a atg gtg gat	963
Ala Lys Asn Leu Gly	Tyr Asn Asp L	ys Tyr Asp Leu Ly	s Met Val Asp	
300	3	05	310	
atg aga aaa att gtg				1011
Met Arg Lys Ile Val	l Pro Glu Thr A	sn Leu Lys Glu Le	eu Ile Lys Asp	
315	320	32	25	
cgt cac tgt gag tc	t gat ttg gac t	gt gtc tat ggc a	ca gat tgt aga	1059
Arg His Cys Glu Se	r Asp Leu Asp (Cys Val Tyr Gly T	hr Asp Cys Arg	
330	335	340		

act agc tgt gat cag agt aca atg aag tgt act to	ca gaa gtg ata caa	1107
Thr Ser Cys Asp Gln Ser Thr Met Lys Cys Thr Se	er Glu Val Ile Gln	
345 350 355	360	
cca aac ttg gca aaa gct tgt cag tta ctc aaa ga	ac tac cta ctg cgt	1155
Pro Asn Leu Ala Lys Ala Cys Gln Leu Leu Lys As	sp Tyr Leu Leu Arg	
365 370	375	
ggt gct cca agt gaa att cgt gaa gaa tta gaa aa	ag cag ctt tat tct	1203
Gly Ala Pro Ser Glu Ile Arg Glu Glu Leu Glu Ly	ys Gln Leu Tyr Ser	
380 385	390	
tgt att gct ctc aaa gtc aca gca aat caa atg ga	aa atg gaa cat tct	1251
Cys Ile Ala Leu Lys Val Thr Ala Asn Gln Met G	lu Met Glu His Ser	
. 395 400	405	
ttg ata cta aat aac cta aaa aca tta ttg tgg a	ag aaa att tcc tac	1299
Leu Ile Leu Asn Asn Leu Lys Thr Leu Leu Trp L	ys Lys Ile Ser Tyr	
410 415 4	120	
act aat gac tot tagttoatt tggacataat taccattt	ta agaaacctgc	1350
Thr Asn Asp Ser		
425		
cacttttaaa gaacaatttt gagcattaaa aaaaaatggc t	ttcaaattcc tgccagttac	1410
acaaaactcc ttcccccag gcctgagaag ccatcagtat g	gtgattactg aagtaatggc	1470
aggtgtagga tcaacaggtc cccaagatgt cattcctgcc c	cttttagaag ccctgttaca	1530
tctccgaagt acattcattg tgtaactatt ttgactgact 1	ttaaaaacca atgctgtgaa	1590
aagcttcatt ccataaacat caacagtgag tgatttgtag a	atttacctta gccaaaatac	1650
caatgctgga agcattgtgt ttgcattgaa gctgctgttc	aacaagaaaa tttataaatt	1710
tectestate ttegestagt easyttiges estimacags	aattaagact gcaaagcagg	1770

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ttaaacttgc	ttctttataa	aacagatgtt	gggttaatag	catggtttac	tgtattaaag	1830
acttatacac	ccatttttaa	cctcattcag	acatcaagtt	atgtgtagct	tcacaatggt	1890
tcaagtggct	tacttcaaga	aatcttatac	ttgacagtac	accaatttta	ttgactaaaa	1950
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ccttcaaaac	tgacatctta	aggcccaatc	aagatccaca	tatcctgatt	ttgaactatg	2070
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catctccata	aaattaaaca	ccttttggag	aaaagatcca	ctattttctg	ctcaaaggtt	2370
togootacot	aaagtggaac	atgttaaaaa	tctatgtgac	catcactgga	cagctttctc	2430
tcaaaacttt	ccttcaacgc	catggattag	caccagtttt	gtttacttta	aggtactttt	2490
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⟨210⟩ 90

<211> 1911

<212> DNA

<213> Homo sapiens

⟨220⟩

<221> CDS

<222> (232)...(1083)

<400> 90

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ttcagagctg tgactgcggc tgcactcaga gaagctgccc ttggctgctc gtagcgccgg 180

gccttc	tctc	ct	cgto	atca	a tc	caga	gcag	cca	gtgto	cg	ggag	gcag	aa g	atg	ccc	237
														Met	Pro	
									•					1		
cac tc	c ag	c c	tg (cat	сса	tcc	atc	ccg	tgt (ccc	agg	ggt	cac	ggg	gcc	285
His Se	r Se	r I	.eu 1	His	Pro	Ser	Ile	Pro	Cys 1	Pro	Arg	G1y	His	Gly	Ala	٠
		5					10					15				
cag aa	ıg go	a g	zcc	ttg	gtt	ctg	ctg	agt	gcc	tgc	ctg	gtg	acc	ctt	tgg	333
Gln Ly	s Al	la I	Ala	Leu	Val	Leu	Leu	Ser	Ala	Cys	Leu	Val	Thr	Leu	Trp	
2	20					25					30					
ggg ct	ta gg	ga į	gag	cca	cca	gag	cac	act	ctc	cgg	tac	ctg	gtg	ctc	cac	381
Gly Le	eu G	ly (Glu	Pro	Pro	Glu	His	Thr	Leu	Arg	Tyr	Leu	Val	Leu	His	
35					40					45					50	
cta g	cc t	сс	ctg	cag	ctg	gga	ctg	ctg	tta	aac	ggg	gtc	tgc	agc	ctg	429
Leu A	la S	er	Leu	Gln	Leu	Gly	Leu	Leu	Leu	Asn	Gly	Val	Cys	Ser	Leu	
				55					60					65		
gct g	ag g	ag	ctg	cac	cac	atc	cac	tcc	agg	tac	cgg	ggo	ago	tac	tgg	477
Ala G	lu G	lu	Leu	His	His	Ile	His	Ser	Arg	Tyr	Arg	Gly	Ser	Tyr	Trp	1
		•	70					75					80)		
agg a	ict g	tg	cgg	gcc	tgc	ctg	ggc	tgc	ccc	ctc	cgc	cgt	ggg	gcc	ctg	525
Arg 1	hr V	al	Arg	Ala	Cys	Leu	Gly	Cys	Pro	Leu	Arg	Are	g G13	, Ala	Leu	1
		85					90)				9	5			
ttg	ctg (ctg	tcc	ato	ta1	t tto	tac	tac	tcc	cto	cca	a aa	t gc	g gto	c ggo	573
Leu l	Leu I	Leu	Ser	· Ile	e Ty	r Phe	э Туз	. Tyı	: Ser	Leu	ı Pro	As	n Ala	a Va	l Gly	y
	100					109	5				110)				
ccg	ccc ·	ttc	act	t tgg	g at	g ct	t gc	c ct	ccte	g gg	cct	c tc	g ca	g gc	a ct	g 621

Pro	Pro	Phe	Thr	Trp	Met	Leu	Ala	Leu	Leu	Gly	Leu	Ser	Gln	Ala	Leu	
115					120					125					130	·
aac	atc	ctc	ctg	ggc	ctc	aag	ggc	ctg	gcc	сса	gct	gag	atc	tct	gca	669
Asn	Ile	Leu	Leu	Gly	Leu	Lys	Gly	Leu	Ala	Pro	Ala	Glu	Ile	Ser	Ala	
				135					140					145		
gtg	tgt	gaa	aaa	ggg	aat	ttc	aac	gtg	gcc	cat	ggg	ctg	gca	tgg	tca	717
Val	Cys	Glu	Lys	Gly	Asn	Phe	Asn	Val	Ala	His	Gly	Leu	Ala	Trp	Ser	
			150					155					160			
tat	tac	atc	gga	tat	ctg	cgg	ctg	atc	ctg	cca	gag	ctc	cag	gcc	cgg	765
Tyr	Tyr	· Ile	Gly	Tyr	Leu	Arg	Leu	Ile	Leu	Pro	Glu	Leu	Gln	Ala	Arg	
		165	;				170					175	•			
att	cga	a act	tac	aat	cag	cat	tac	aac	aac	ctg	cta	cgg	ggt	gca	gtg	813
Ile	Arg	g Thr	Tyı	. Asr	Gln	His	Tyr	Asn	Asn	Leu	Leu	Arg	Gly	, Ala	a Val	
	180	0				185	;				190)				
ago	ca	g cg	g cta	g tai	tatt	ctc	ctc	cca	a ttg	gac	tgt	ggs	g gtg	g cc	t gat	861
Sea	r Gl	n Arı	g Lei	и Туз	r Ile	e Leu	. Leu	Pro	Leu	. Asp	Cys	G1;	y Val	l Pr	o Asp	
19	5				200)				205					210	
aa	c ct	g ag	t at	g gc	t gad	c ccc	c aac	at	t cgc	tto	ct	g ga	t aa	a ct	g ccc	909
As	n Le	u Se	r Me	t Al	a Ası	p Pro	o Ası	ı Il	e Arg	g Phe	Lei	ı As	p Ly	s Le	u Pro	
				21	5				220)				22	5	
ca	g ca	g ac	c gc	t ga	c cg	t gc	t gg	c at	c aa	g gat	t cg	g gt	t ta	c ag	c aac	957
G1	n Gl	n Th	r Al	a As	p Ar	g Ala	a Gl	y Il	e Ly	s Ası	Ar	g Va	l Ty	r Se	r Asn	•
			23	80				23	5				24	10		
ag	c at	tc ta	at ga	ag ct	t ct	g ga	g aa	c gg	g ca	g cg	g aa	c ct	g ca	ig at	g aca	1005
S	~ T1	ד בו	,- G1	lu Le	eu Le	u Gl	u As	n G1	y G1	n Ar	g As	n Le	eu Gl	ln Me	et Thr	•

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245	250	255	
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Ala Ala Ser Arg Cys Pro		•	
260	265	270	
aaa agg aag agg tta ctg	tgg gca gct tg	aagacctc agcggtgccc	1100
Lys Arg Lys Arg Leu Leu			
275 280			•
agtacctcca cgatgtccca a	ngagootgag otoot	catca gtggaatgga aaa	agcccctc 1160
cctctccgca cggatttctc 1			
ccaagcetet ggactggggg			
ccacaggggg ccttgcaggg			
cttgggccag tcatttcccc			
taatcactgc cttacctccc			
taaactttgg atgctagtgt			
gacccactcc ccacccttct			
atcaacaggc tccttcgccc	tctggctcct ggtc	atgttc cattattggg ga	gccccagc 1640
agaagaatgg agaggaggag	gaggctgagt ttgg	ggtatt gaatcccccg gc	tcccaccc 1700
tgcagcatca aggttgctat			
ctctaggatt ctggcaccac	•		
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<210> 91

<211> 476

<212> PRT

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(400)	> 91														
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1				5					10					15	
Pro	Gly	Pro	Cys	Asp	Gly	Leu	Phe	Arg	Ser	Leu	Tyr	Arg	Ser	Val	Ser
			20					25					30		
Met	Pro	Pro	Lys	Gly	Asp	Ser	Gly	Gln	Pro	Leu	Phe	Leu	Thr	Pro	Tyr
		35					40					45			
Ile	Glu	Ala	Gly	Lys	Ile	Gln	Lys	Gly	Arg	Glu	Leu	Ser	Leu	Val	Gly
	50					55					60				
Pro	Phe	Pro	Gly	Leu	Asn	Met	Lys	Ser	Tyr	Ala	Gly	Phe	Leu	Thr	Val
65					70				•	75					80
Asn	Lys	Thr	Tyr	Asn	Ser	Asn	Leu	Phe	Phe	Trp	Phe	Phe	Pro	Ala	Gln
				85					90					95	
Ile	Gln	Pro	Glu	Asp	Ala	Pro	Val	Val	Leu	Trp	Leu	Gln	Gly	Gly	Pro
			100)				105					110		
Gly	Gly	Ser	Ser	Met	Phe	Gly	Leu	Phe	Val	Glu	His	Gly	Pro	Tyr	Val
		115	;				120					125	5		
Val	Thr	Ser	Asr	Met	Thr	Leu	Arg	Asp	Arg	Asp	Phe	Pro	Trp	Thr	Thr
	130)				135	I				140)			
Thr	Leu	Ser	Me1	t Leu	ı Tyr	· Ile	Asp	Asn	Pro	Val	Gly	Thi	c Gly	Phe	Ser
145	;				150)				155	5				160
Phe	Thi	r Ası	Ası	p Thi	r His	Gly	Tyr	Ala	Val	l Asr	ı Glu	ı Ası	p Ası	va]	l Ala
				169	5				170)				179	5
Arg	, Ası	Lei	ע Ty:	r Se	r Ala	a Lei	ı Ile	e Gl	n Pho	e Phe	e Gli	n II	e Ph	e Pro	o Glu

			180					185					190		
Tyr	Lys	Asn	Asn	Asp	Phe	Tyr	Val	Thr	Gly (Glu	Ser '	Tyr	Ala	Gly	Lys
		195					200	•				205			
Tyr	Val	Pro	Ala	Ile	Ala	His	Leu	Ile	His	Ser	Leu	Asn	Pro	Val	Arg
	210					215					220				
Glu	Val	Lys	Ile	Asn	Leu	Asn	Gly	Ile	Ala	Ile	Gly	Asp	Gly	Tyr	Ser
225					230					235					240
Asp	Pro	Glu	Ser	Ile	Ile	Gly	Gly	Tyr	Ala	Glu	Phe	Leu	Tyr	Gln	Ile
				245					250					255	
G1y	Leu	Leu	Asp	Glu	Lys	Gln	Lys	Lys	Tyr	Phe	Gln	Lys	Gln	Cys	His
			260					265					270		
Glu	Cys	Ile	Glu	His	Ile	Arg	Lys	Gln	Asn	Trp	Phe	Glu	Ala	Phe	Glu
		275					280					285			
Ile	Leu	. Asp	Lys	Leu	Leu	Asp	Gly	Asp	Leu	Thr	Ser	Asp	Pro	Ser	Tyr
	290)			٠	295					300				
Phe	Glr	ı Asn	val	. Thr	Gly	Cys	Ser	Asn	Tyr	Tyr	Asn	Phe	Leu	Arg	Cys
305	;				310)				315					320
Thr	Glu	ı Pro	Glu	ı Asp	Glr	Leu	Tyr	Tyr	Val	Lys	Phe	Leu	Ser	Leu	Pro
		ė		325	5				330)				335	5
Glu	ı Va	l Ara	g Glı	n Ala	a Ile	His	. Val	Gly	Asn	Gln	Thr	Phe	. Asr	ı Asp	Gly
			340	0				345	j				350)	
Thi	· 11	e Va	l Gl	u Ly:	s Ty	r Lei	ı Arg	g Glu	ı Asp	Th:	· Val	Glr	ı Sei	r Val	l Lys
		35	5				360)				369	5		
Pre	o Tr	p Le	u Th	r Gl	u Il	e Me	t Ası	n Asr	ı Tyı	r Ly:	s Val	l Lei	u Il	е Ту	r Asn
	37	0				37	5				380	0			

197/307

Gly Gln Leu Asp Ile Ile Val Ala Ala Ala Leu Thr Glu His Ser Leu Met Gly Met Asp Trp Lys Gly Ser Gln Glu Tyr Lys Lys Ala Glu Lys Lys Val Trp Lys Ile Phe Lys Ser Asp Ser Glu Val Ala Gly Tyr Ile Arg Gln Ala Gly Asp Phe His Gln Val Ile Ile Arg Gly Gly His Ile Leu Pro Tyr Asp Gln Pro Leu Arg Ala Phe Asp Met Ile Asn Arg Phe Ile Tyr Gly Lys Gly Trp Asp Pro Tyr Val Gly <210> 92 <211> 226 <212> PRT <213> Homo sapiens <400> 92 Met Ser Arg Ala Gln Ile Trp Ala Leu Val Ser Gly Val Gly Gly Phe Gly Ala Leu Val Ala Ala Thr Thr Ser Asn Glu Trp Lys Val Thr Thr Arg Ala Ser Ser Val Ile Thr Ala Thr Trp Val Tyr Gln Gly Leu Trp Met Asn Cys Ala Gly Asn Ala Leu Gly Ser Phe His Cys Arg Pro His

	50					55					60				
Phe	Thr	Ile	Phe	Lys	Val	Ala	Gly	Tyr	Ile	Gln	Ala	Cys	Arg	Gly	Leu
65		•			70					75					80
Met	Ile	Ala	Ala	Val	Ser	Leu	Gly	Phe	Phe	Gly	Ser	Ile	Phe	Ala	Leu
				85					90					95	
Phe	Gly	Met	Lys	Cys	Thr	Lys	Val	Gly	Gly	Ser	Asp	Lys	Ala	Lys	Ala
			100					105					110		
Lys	Ile	Ala	Cys	Leu	Ala	Gly	Ile	Val	Phe	Ile	Leu	Ser	Gly	Leu	Cys
		115	,				120					125			
Ser	Met	Thr	Gly	Cys	Ser	Leu	Tyr	Ala	Asn	Lys	Ile	Thr	Thr	Glu	Phe
	130)				135					140				
Phe	Asp	Pro	Lev	ı Phe	· Val	Glu	G1n	Lys	Tyr	Glu	l Leu	Gly	Ala	Ala	Leu
145	i				150)				155	5 .				160
Phe	: Ile	e Gly	y Trj	o Ala	a Gly	, Ala	s Ser	Leu	ı Cys	s Ile	e Ile	Gly	y Gly	Val	Ile
				165	5				170)				175	i
Phe	e Cy	s Ph	e Se	r Ile	e Se	r Ası	Ası	n Ası	n Lys	s Th	r Pro	Ar	g Tyı	Thr	Tyr
			18	0				18	5				190)	
Ası	n Gl	y Al	a Th	r Se	r Va	l Me	t Se	r Se	r Ar	g Th	r Ly	s Ty	r Hi	s Gly	Gly
-		19	5				20	0				20	5		
G1	u As	p Ph	e Ly	s Th	r Th	r As	n Pr	o Se	r Ly	s Gl	n Ph	e As	p Ly	s Ası	n Ala
	21	0				21	5				22	0			
Ту	r Va	1													
22	5														

2211> 305
212> PRT
(213> Homo sapiens
<400> 93
Met Gly Ile Gln Thr Ser Pro Val Leu Leu Ala Ser Leu Gly Val Gly
1 5 10 15
Leu Val Thr Leu Leu Gly Leu Ala Val Gly Ser Tyr Leu Val Arg Arg
20 25 30
Ser Arg Arg Pro Gln Val Thr Leu Leu Asp Pro Asn Glu Lys Tyr Leu
35 40 45
Leu Arg Leu Leu Asp Lys Thr Thr Val Ser His Asn Thr Lys Arg Phe
50 55 60
Arg Phe Ala Leu Pro Thr Ala His His Thr Leu Gly Leu Pro Val Gly
65 70 75 80
Lys His Ile Tyr Leu Ser Thr Arg Ile Asp Gly Ser Leu Val Ile Arg
85 90 95
Pro Tyr Thr Pro Val Thr Ser Asp Glu Asp Gln Gly Tyr Val Asp Leu
100 105 110
Val Ile Lys Val Tyr Leu Lys Gly Val His Pro Lys Phe Pro Glu Gly
115 120 125
Gly Lys Met Ser Gln Tyr Leu Asp Ser Leu Lys Val Gly Asp Val Val
130 135 140
Glu Phe Arg Gly Pro Ser Gly Leu Leu Thr Tyr Thr Gly Lys Gly His
145 150 155 160
Phe Asn Ile Gln Pro Asn Lys Lys Ser Pro Pro Glu Pro Arg Val Ala

				165					170					175	
Lys	Lys	Leu	Gly	Met	Ile	Ala	Gly	Gly	Thr	Gly	Ile	Thr	Pro	Met	Leu
			180					185					190		
Gln	Leu	Ile	Arg	Ala	Ile	Leu	Lys	Val	Pro	Glu	Asp	Pro	Thr	Gln	Cys
		195					200					205			
Phe	Leu	Leu	Phe	Ala	Asn	Gln	Thr	Glu	Lys	Asp	Ile	Ile	Leu	Arg	Glu
	210					215					220				
Asp	Leu	Glu	Glu	Leu	Gln	Ala	Arg	Tyr	Pro	Asn	Arg	Phe	Lys	Leu	Trp
225					230					235					240
Phe	Thr	Leu	Asp	His	Pro	Pro	Lys	Asp	Trp	Ala	Tyr	Ser	Lys	Gly	Phe
				245					250					255	
Val	Thr	Ala	Asp	Met	Ile	Arg	Glu	His	Leu	Pro	Ala	Pro	Gly	Asp	Asp
			260	•				265					270		
Val	Leu	ı Val	Leu	Leu	Cys	Gly	Pro	Pro	Pro	Met	Val	Gln	Leu	Ala	Cys
		275	;				280					285			
His	Pro	Asr	Leu	Asp	Lys	Leu	Gly	Tyr	Ser	Gln	Lys	Met	Arg	Phe	Thr
	290)				295	;				300				
Tyr	•														
305	5														
<2 1	10>	94													
<2	1>	227													
<2 :	12>	PRT													
<2	13>	Ното	sap	iens											
< 4	00>	94													

Met	Gly	Trp	Thr	Met	Arg	Leu	Val	Thr	Ala	Ala	Leu	Leu	Leu	Gly	Leu
. 1				5					10			٠		15	
Met	Met	Val	Val	Thr	Gly	Asp	Glu	Asp	Glu	Asn	Ser	Pro	Cys	Ala	His
			20					25					30		
Glu	Ala	Leu	Leu	Asp	Glu	Asp	Thr	Leu	Phe	Cys	Gln	Gly	Leu	Glu	Val
		35					40					45			
Phe	Tyr	Pro	Glu	Leu	Gly	Asn	Ile	Gly	Cys	Lys	Val	Val	Pro	Asp	Cys
	50					55					60				
Asn	Asn	Tyr	Arg	Gln	Lys	Ile	Thr	Ser	Trp	Met	Glu	Pro	Ile	Val	Lys
65					70					75					80
Phe	Pro	Gly	Ala	Val	Asp	Gly	Ala	Thr	Tyr	Ile	Leu	Val	Met	Val	Asp
				85					90					95	
Pro	Asp	Ala	Pro	Ser	Arg	Ala	Glu	Pro	Arg	Gln	Arg	Phe	Trp	Arg	His
			100					105					110		
Trp	Leu	Val	Thr	Asp	Ile	Lys	Gly	Ala	Asp	Leu	Lys	Lys	Gly	Lys	Ile
		115	i				120)				125	•		
Gln	Gly	Gln	Glu	Leu	Ser	Ala	Tyr	Gln	Ala	Pro	Ser	Pro	Pro	Ala	His
	130)				135)				140)			
Ser	Gly	Phe	His	Arg	Tyr	Gln	Phe	Phe	Val	Tyr	Leu	Glr	ı Glu	Gly	Lys
145	;				150)				155	,				160
Val	Ile	Ser	Leu	Leu	Pro	Lys	Glu	ı Asn	Lys	Thr	Arg	Gly	/ Ser	Trp	Lys
				165	,				170)				175	i
Met	. Asp	Are	g Phe	. Leu	ı Asr	Arg	, Phe	His	Leu	Gly	/ Glu	ı Pro	o Glu	Ala	Ser
			180)				185	,				190	}	
Th	. C1.	. Dh	. Not	The	· Glr	Asr	. Tvi	- Glr	Asr	s Sei	r Pro	Thi	r Leu	Glr	Ala

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Pro Arg Glu Arg Ala Ser Glu Pro Lys His Lys Asn Gln Ala Glu Ile 210 . Ala Ala Cys <210> 95 <211> 441 <212> PRT <213> Homo sapiens <400> 95 Met Ala Ile His Lys Ala Leu Val Met Cys Leu Gly Leu Pro Leu Phe Leu Phe Pro Gly Ala Trp Ala Gln Gly His Val Pro Pro Gly Cys Ser Gln Gly Leu Asn Pro Leu Tyr Tyr Asn Leu Cys Asp Arg Ser Gly Ala Trp Gly Ile Val Leu Glu Ala Val Ala Gly Ala Gly Ile Val Thr Thr Phe Val Leu Thr Ile Ile Leu Val Ala Ser Leu Pro Phe Val Gln Asp Thr Lys Lys Arg Ser Leu Leu Gly Thr Gln Val Phe Phe Leu Leu Gly Thr Leu Gly Leu Phe Cys Leu Val Phe Ala Cys Val Val Lys Pro Asp

he	Ser	Thr	Cys	Ala	Ser	Arg	Arg	Phe	Leu	Phe	Gly	Val	Leu	Phe	Ala	
		115					120					125				
[]e	Cys	Phe	Ser	Cys	Leu	Ala	Ala	His	Val	Phe	Ala	Leu	Asn	Phe	Leu	
	130					135					140					
Ala	Arg	Lys	Asn	His	Gly	Pro	Arg	Gly	Trp	Val	Ile	Phe	Thr	Val	Ala	
145					150					155					160	
Leu	Leu	Leu	Thr	Leu	Val	Glu	Val	.Ile	Ile	Asn	Thr	Glu	Trp	Leu	Ile	
				165					170					175		
Ile	Thr	Leu	Val	Arg	Gly	Ser	Gly	Glu	Gly	Gly	Pro	Gln	Gly	Asn	Ser	
			180					185					190)		
Ser	Ala	Gly	Trp	Ala	Val	Ala	Ser	Pro	Cys	Ala	Ile	Ala	Asn	Met	Asp	
		195					200			•		205	j			
Phe	Val	Met	Λla	Leu	Ile	Tyr	Val	Met	Leu	Leu	Leu	Lec	ı Gly	Ala	Phe	
	210	,				215					220)				
Leu	Gly	Ala	Trp	Pro	Ala	Leu	Cys	Gly	Arg	Tyr	Lys	: Ar	g Trp	Arg	g Lys	
225					230					235	;				240	
His	Gly	Va]	Phe	e Val	Leu	Leu	Thr	Thr	Ala	Thr	Sei	· Va.	l Ala	a Ile	Trp	
				245	,				250)				25	5	
Val	Val	Tr	lle	e Val	. Met	Tyr	The	Tyr	Gly	/ Asr	ı Lys	s Gl	n Hi	s Ası	n Ser	
			260)				265	5				27	0		
Pro	Thi	r Trj	a Ası	o Asp) Pro	Thr	Let	ı Ala	ılle	e Ala	a Le	u Al	a Al	a Ası	n Ala	
		27	5				280)		•		28	5			
Trp	Ala	a Pho	e Val	l Lei	ı Phe	у Туз	· Vai	l I1e	e Pro	o Gli	u Va	l Se	r Gl	n Va	1 Thr	
	290	0				299	5				30	0				
1.ve	s Sei	r Se	r Pr	o Gli	ı Glr	ı Sei	r Ty:	r Gli	n Gl	y As	p Me	t Ty	r Pr	o Th	r Arg	

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Gly Val Gly Tyr Glu Thr Ile Leu Lys Glu Gln Lys Gly Gln Ser Met Phe Val Glu Asn Lys Ala Phe Ser Met Asp Glu Pro Val Ala Ala Lys Arg Pro Val Ser Pro Tyr Ser Gly Tyr Asn Gly Gln Leu Leu Thr Ser Val Tyr Gln Pro Thr Glu Met Ala Leu Met His Lys Val Pro Ser Glu Gly Ala Tyr Asp Ile Ile Leu Pro Arg Ala Thr Ala Asn Ser Gln Val Met Gly Ser Ala Asn Ser Thr Leu Arg Ala Glu Asp Met Tyr Ser Ala Gln Ser His Gln Ala Ala Thr Pro Pro Lys Asp Gly Lys Asn Ser Gln Val Phe Arg Asn Pro Tyr Val Trp Asp <210> 96 <211> 265 <212> PRT <213> Homo sapiens <400> 96 Met Ala Ala Ala Val Pro Lys Arg Met Arg Gly Pro Ala Gln Ala Lys

l

Leu	Leu	Pro	Gly	Ser	Ala	Ile	Gln	Ala	Leu	Val	Gly	Leu	Ala	Arg	Pro
			20				•	25					30		
Leu	Val	Leu	Ala	Leu	Leu	Leu	Val	Ser	Ala	Ala	Leu	Ser	Ser	Val	Val
		35					40					45			
Ser	Arg	Thr	Asp	Ser	Pro	Ser	Pro	Thr	Val	Leu	Asn	Ser	His	Ile	Ser
	50					55					60				
Thr	Pro	Asn	Val	Asn	Ala	Leu	Thr	His	Glu	Asn	Gln	Thr	Lys	Pro	Ser
65					70					75					80
Ile	Ser	Gln	Ile	Ser	Thr	Thr	Leu	Pro	Pro	Thr	Thr	Ser	Thr	Lys	Lys
				85					90					95	
Ser	Gly	Gly	Ala	Ser	Val	Val	Pro	His	Pro	Ser	Pro	Thr	Pro	Leu	Ser
			100					105					110		
Gln	Glu	Glu	ıAla	Asp	Asn	Asn	Glu	Asp	Pro	Ser	Ile	Glu	Glu	Glu	Asp
		115	;				120					125	;		
Leu	Leu	Met	Leu	Asn	Ser	Ser	Pro	Ser	Thr	Ala	Lys	Asp	Thr	Leu	Asp
	130	1				135	;				140)			
Asn	Gly	Asp) Tyr	Gly	Glu	Pro	Asp	Tyr	Asp	Trp	Thr	Thr	Gly	Pro	Arg
145	;				150					155	5				160
Asp	Asp	Asp	o Glu	Ser	Asp	Asp	Thr	Leu	Glu	Glu	ı Asr	n Arg	g Gly	Tyr	Met
				165	i				170)				175	i
Glu	Ile	Glu	ı Glm	Ser	Val	Lys	s Ser	Phe	Lys	s Met	t Pro	Sei	r Sei	Asr	Ile
			180)				189	5				190)	
Glu	ı Glu	ı Glu	u Asp	Ser	His	Phe	e Phe	Phe	His	s Lei	u Ile	e Ile	e Phe	e Ala	. Phe
		19	5				200)				20	5		
Cvs	s Ile	e Ala	a Val	l Val	Tyr	· 11e	e Thr	Tyı	r His	s Ası	n Ly:	s Ar	g Ly:	s Ile	Phe

PCT/JP00/05356 WO 01/12660

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Leu Leu Val Gln Ser Arg Lys Trp Arg Asp Gly Leu Cys Ser Lys Thr Val Glu Tyr His Arg Leu Asp Gln Asn Val Asn Glu Ala Met Pro Ser Leu Lys Ile Thr Asn Asp Tyr Ile Phe <210> 97 <211> 208 <212> PRT <213> Homo sapiens <400> 97 Met Leu Gly Leu Leu Val Ala Leu Leu Ala Leu Gly Leu Ala Val Phe ı Ala Leu Leu Asp Val Trp Tyr Leu Val Arg Leu Pro Cys Ala Val Leu Arg Ala Arg Leu Leu Gln Pro Arg Val Arg Asp Leu Leu Ala Glu Gln Arg Phe Pro Gly Arg Val Leu Pro Ser Asp Leu Asp Leu Leu Leu His Met Asn Asn Ala Arg Tyr Leu Arg Glu Ala Asp Phe Ala Arg Val Ala His Leu Thr Arg Cys Gly Val Leu Gly Ala Leu Arg Glu Leu Arg Ala

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His Thr Val Leu Ala Ala Ser Cys Ala Arg His Arg Arg Ser Leu Arg Leu Leu Glu Pro Phe Glu Val Arg Thr Arg Leu Leu Gly Trp Asp Asp Arg Ala Phe Tyr Leu Glu Ala Arg Phe Val Ser Leu Arg Asp Gly Phe Val Cys Ala Leu Leu Arg Phe Arg Gln His Leu Leu Gly Thr Ser Pro Glu Arg Val Val Gln His Leu Cys Gln Arg Arg Val Glu Pro Pro Glu Leu Pro Ala Asp Leu Gln His Trp Ile Ser Tyr Asn Glu Ala Ser Ser Gln Leu Leu Arg Met Glu Ser Gly Leu Ser Asp Val Thr Lys Asp Gln <210> 98 <211> 400 <212> PRT <213> Homo sapiens <400> 98 Met Ala Trp Arg Arg Glu Ala Ser Val Gly Ala Arg Gly Val Leu Ala Leu Ala Leu Leu Ala Leu Ala Leu Cys Val Pro Gly Ala Arg Gly Arg Ala Leu Glu Trp Phe Ser Ala Val Val Asn Ile Glu Tyr Val Asp

		35					40					45			
Pro C	Gln	Thr	Asn	Leu	Thr	Val	Trp	Ser	Val	Ser	Glu	Ser	Gly	Arg	Phe
	50					55			•	•	60		•		
Gly A	Asp	Ser	Ser	Pro	Lys	Glu	Gly	Ala	His	Gly	Leu	Val	Gly	Val	Pro
65					70					75					80
Trp /	Ala	Pro	Gly	Gly	Asp	Leu	Glu	Gly	Cys	Ala	Pro	Asp	Thr	Arg	Phe
•				85					90					95	
Phe '	Val	Pro	Glu		Gly	Gly	Arg	Gly	Ala	Ala	Pro	Trp	Val	Ala	Leu
			100					105					110		
Val	Ala	Arg			Cys	Thr	Phe	Lys	Asp	Lys	Val	Leu	Val	Ala	Ala
		115					120					125			
Arg	Arg			Ser	Ala	Val	Val	Leu	.Tyr	· Asn	Glu	Glu	Arg	Tyr	Gly
	130					135					140				
Acn		Thr	· f.eu	. Pro) Met			Ala	Gly	, Thr	Gly	Asn	Ile	Val	Val
145	110		500		150				·	155					160
	Mat	Tle	. Sai	- Tur			: G1 v	, Are	Glu			ı Glu	ı Leu	Val	Gln
116	WEL	116	361	169		, 5,5	,	6	170					175	
	C1	. 71.	. D			· Mat	Thr	- Ila			l G1v	, Thr	· Arg		. Val
Lys	огу	116			L 1111	nie t		185		,			190		
0.1	C1	. Di	180		. C1.	. C1.	. Sa			l Ph	⊳ Va`	1 A1:			. Phe
GIN	GIU			e Se	r ui)	011			, , ,			209			
		19				_	200				. 71			а Т у у	- 110
Ile	Thr	Me	t Me	t II	e lle			u Ala	a ir	р ге			e iyi	. iyi	: Ile
	210					21					22			. 112	_ A
Gln	Arg	g Ph	e Le	и Ту	r Th	r Gl	y Se	r Gl	n Il			r Gl	n Sei	r Hi:	s Arg
225					23	0				23	5				240

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Lys	Glu	Thr	Lys	Lys	Val	Ile	Gly	Gln	Leu	Leu	Leu	His	Thr	Val	Lys
				245	•				250					255	
His	Gly	Glu	Lys	Gly	Ile	Asp	Val	Asp	Ala	Glu	Asn	Cys	Ala	Val	Cys
			260					265					270		
Ile	Glu	Asn	Phe	Lys	Val	Lys	Asp	Ile	Ile	Arg	Ile	Leu	Pro	Cys	Lys
		275					280					285			
His	Ile	Phe	His	Arg	Ile	Cys	Ile	Asp	Pro	Trp	Leu	Leu	Asp	His	Arg
	290					295					300				
Thr	Cys	Pro	Met	Cys	Lys	Leu	Asp	Val	Ile	Lys	Ala	Leu	Gly	Tyr	Trp
305					310					315					320
Gly	Glu	Pro	Gly	Asp	Val	Gln	Glu	Met	Pro	Ala	Pro	Glu	Ser	Pro	Pro
				325	1				330	1				335	
Gly	Arg	Asp	Pro	Ala	Ala	Asn	Leu	Ser	Leu	Ala	Leu	Pro	Asp	Asp	Asp
			340)				345	;				350		
Gly	/ Ser	· Asp	Glu	. Ser	Ser	Pro	Pro	Ser	Ala	Ser	Pro	Λla	Glu	Ser	Glu
		359	5				360)				365	;		
Pro	Glr	Cys	s Asp	Pro	Ser	Phe	e Lys	Gly	/ Asp	Ala	Gly	Glu	ı Asn	Thr	· Ala
	370)				379	5				380)			
Lei	ı Lev	ı Glu	ı Ala	a Gly	/ Arg	Sei	. Asp	Sei	r Arg	g His	Gly	Gly	/ Pro	Ile	e Ser
38	5				390)				395	5				400

⟨210⟩ 99

<211> 192

<212> PRT

<213> Homo sapiens

(400)	> 99)													
Met 1	Phe	Cys	Pro	Leu	Lys	Leu	Ile	Leu	Leu	Pro	Val	Leu	Leu	Asp	Tyr
1				5	•				10					15	
Ser	Leu	Gly	Leu	Asn	Asp	Leu	Asn	Val	Ser	Pro	Pro	Glu	Leu	Thr	Val
			20					25					30		
His	Val	Gly	Asp	Ser	Ala	Leu	Met	Gly	Cys	Val	Phe	Gln	Ser	Thr	Glu
		35					40					45			
Asp	Lys	Cys	Ile	Phe	Lys	Ile	Asp	Trp	Thr	Leu	Ser	Pro	Gly	Glu	His
	50					55					60				
Ala	Lys	Λsp	Glu	Tyr	Val	Leu	Tyr	Tyr	Tyr	Ser	Asn	Leu	Ser	Val	Pro
65					70					75					80
Ile	Gly	۸re	g Phe	Gln	Asn	Arg	Val	His	Leu	Met	Gly	Asp	Asn	Leu	Cys
				85	;				90)	•			95	i
Asn	Asp	Gly	y Sei	r Leu	ı Leu	ı Let	Gln	Asp	Val	Glr	Glu	Ala	Asp	Gln	Gly
			100)			•	105	5				110)	
Thr	Туі	c Ile	e Cy:	s Glu	ı Ile	e Are	g Leu	Lys	s Gly	y Glu	ı Ser	Glr	ı Val	l Phe	e Lys
		11	5				120)				129	5		
Lys	Ala	a Va	l Va	l Lei	u Hi:	s Va	l Lei	ı Pro	o Gl	u Glı	ı Pro	Lys	s Glu	ı Lei	ı Met
	13	0		·		13	5				140)			
Val	Hi	s Va	1 G1	y Gl	y Le	u Il	e Glı	n Me	t Gl	у Су	s Va	l Ph	e Gl	n Se	r Thr
145	5				15	0				15	5				160
Glu	ı Va	l Ly	s Hi	s Va	l Th	r Ly	s Va	1 G1	u Tr	p Il	e Ph	e Se	r Gl	y Ar	g Arg
				16	5				17	0				17	5
Ala	a Ly	s Va	ıl Th	ır Ar	g Ar	g Ly	s Hi	s Hi	s Cy	's Va	1 Ar	g Gl	u Gl	y Se	r Gly
			18	30				18	5				19	0	

> 10	0													
> 26	60													•
?> PF	RT													
3> Ho	omo s	apie	ens											
)> 10	00													
Ala	Gly	Ser	Pro	Leu	Leu	Trp	Gly	Pro	Arg	Ala	Gly	Gly	Val	Gly
			5					10					15	
Leu	Val	Leu	Leu	Leu	Leu	Gly	Leu	Phe	Arg	Pro	Pro	Pro	Ala	Leu
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Ala	Arg	Pro	Val	Lys	Glu	Pro	Arg	Gly	Leu	Ser	Ala	Ala	Ser	Pro
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Leu	Leu	Glu	Ala	Glu	Arg	Gln	Glu	Arg	Ala	Arg	Ala	Glu	Ala	Gln
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130					135					140				
		Ala	Ala	Leu	Ala	Ala	Gln	Leu	Val	Pro	Ala	Pro	Val	Pro
	> 26 PP PR S Ho Ala Leu Ala Leu Ala Gly Leu Ala 130	D> 100 Ala Gly Leu Val Ala Arg 35 Leu Ala 50 Gly Glu Leu Leu Ala Glu Gly Ala 115 Asp Ala 130	> 260 PRT S Homo sapie 10 100 Ala Gly Ser Leu Val Leu 20 Ala Arg Pro 35 Leu Ala Glu 50 Gly Glu Ala Leu Leu Glu Ala Glu Asp 100 Gly Ala Pro 115 Asp Ala Pro 130	> 260 PRT S Homo sapiens 100 Ala Gly Ser Pro 5 Leu Val Leu Leu 20 Ala Arg Pro Val 35 Leu Ala Glu Thr 50 Gly Glu Ala Ala Leu Leu Glu Ala 85 Ala Glu Asp Gln 100 Gly Ala Pro Arg 115 Asp Ala Pro Ala	> 260 2> PRT 3> Homo sapiens 3> 100 Ala Gly Ser Pro Leu	> 260 PRT S Homo sapiens 100 Ala Gly Ser Pro Leu Leu 5 Leu Val Leu Leu Leu Leu 20 Ala Arg Pro Val Lys Glu 35 Leu Ala Glu Thr Gly Ala 50 55 Gly Glu Ala Ala Gly Ala 70 Leu Leu Glu Ala Glu Arg 85 Ala Glu Asp Gln Gln Ala 100 Gly Ala Pro Arg Asn Ser 115 Asp Ala Pro Ala Ala Gln 130 135	> 260 2> PRT 3> Homo sapiens 3> 100 Ala Gly Ser Pro Leu Leu Trp 5 Leu Val Leu Leu Leu Leu Gly 20 Ala Arg Pro Val Lys Glu Pro 35 40 Leu Ala Glu Thr Gly Ala Pro 50 55 Gly Glu Ala Ala Gly Ala Val 70 Leu Leu Glu Ala Glu Arg Gln 85 Ala Glu Asp Gln Gln Ala Arg 100 Gly Ala Pro Arg Asn Ser Asp 115 120 Asp Ala Pro Ala Ala Gln Leu 130 135	> 260 >> PRT S Homo sapiens >> 100 Ala Gly Ser Pro Leu Leu Trp Gly 5 Leu Val Leu Leu Leu Leu Gly Leu 20 25 Ala Arg Pro Val Lys Glu Pro Arg 35 40 Leu Ala Glu Thr Gly Ala Pro Arg 50 55 Gly Glu Ala Ala Gly Ala Val Gln 70 Leu Leu Glu Ala Glu Arg Gln Glu 85 Ala Glu Asp Gln Gln Ala Arg Val 100 105 Gly Ala Pro Arg Asn Ser Asp Pro 115 120 Asp Ala Pro Ala Ala Gln Leu Ala 130 135	> 260 2> PRT 3> Homo sapiens 3> 100 Ala Gly Ser Pro Leu Leu Trp Gly Pro	> 260 >> PRT S Homo sapiens > 260 2> PRT 3> Homo sapiens 3> 100 Ala Gly Ser Pro Leu Leu Trp Gly Pro Arg Ala 5 10 Leu Val Leu Leu Leu Leu Gly Leu Phe Arg Pro 20 25 Ala Arg Pro Val Lys Glu Pro Arg Gly Leu Ser 35 40 Leu Ala Glu Thr Gly Ala Pro Arg Arg Phe Arg 50 55 60 Gly Glu Ala Ala Gly Ala Val Gln Glu Leu Ala 70 75 Leu Leu Glu Ala Glu Arg Gln Glu Arg Ala Arg 85 90 Ala Glu Asp Gln Gln Ala Arg Val Leu Ala Gln 100 105 Gly Ala Pro Arg Asn Ser Asp Pro Ala Leu Gly 115 120 Asp Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu 130 135 140	> 260 2> PRT 3> Homo sapiens 3> 100 Ala Gly Ser Pro Leu Leu Trp Gly Pro Arg Ala Gly 5 10 Leu Val Leu Leu Leu Leu Gly Leu Phe Arg Pro Pro 20 25 Ala Arg Pro Val Lys Glu Pro Arg Gly Leu Ser Ala 35 40 45 Leu Ala Glu Thr Gly Ala Pro Arg Arg Phe Arg Arg 50 55 60 Gly Glu Ala Ala Gly Ala Val Gln Glu Leu Ala Arg 70 75 Leu Leu Glu Ala Glu Arg Gln Glu Arg Ala Arg Ala 85 90 Ala Glu Asp Gln Gln Ala Arg Val Leu Ala Gln Leu 100 105 Gly Ala Pro Arg Asn Ser Asp Pro Ala Leu Gly Leu 115 120 125 Asp Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu Leu 130 135 140	> 260 2> PRT 3> Homo sapiens 3> 100 Ala Gly Ser Pro Leu Leu Trp Gly Pro Arg Ala Gly Gly 5 10 Leu Val Leu Leu Leu Leu Gly Leu Phe Arg Pro Pro Pro 20 25 30 Ala Arg Pro Val Lys Glu Pro Arg Gly Leu Ser Ala Ala 35 40 45 Leu Ala Glu Thr Gly Ala Pro Arg Arg Phe Arg Arg Ser 50 55 60 Gly Glu Ala Ala Gly Ala Val Gln Glu Leu Ala Arg Ala 70 75 Leu Leu Glu Ala Glu Arg Gln Glu Arg Ala Arg Ala Glu 85 90 Ala Glu Asp Gln Gln Ala Arg Val Leu Ala Gln Leu Leu 100 105 110 Gly Ala Pro Arg Asn Ser Asp Pro Ala Leu Gly Leu Asp 115 120 125 Asp Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu Leu Arg 130 135 140	> 260 >> PRT >> Homo sapiens >> 100 Ala Gly Ser Pro Leu Leu Trp Gly Pro Arg Ala Gly Gly Val 5 10 15 Leu Val Leu Leu Leu Leu Gly Leu Phe Arg Pro Pro Pro Ala 20 25 30 Ala Arg Pro Val Lys Glu Pro Arg Gly Leu Ser Ala Ala Ser 35 40 45 Leu Ala Glu Thr Gly Ala Pro Arg Arg Phe Arg Arg Ser Val 50 55 60 Gly Glu Ala Ala Gly Ala Val Gln Glu Leu Ala Arg Ala Leu 70 75 Leu Leu Glu Ala Glu Arg Gln Glu Arg Ala Arg Ala Glu Ala 85 90 95 Ala Glu Asp Gln Gln Ala Arg Val Leu Ala Gln Leu Leu Arg 100 105 110 Gly Ala Pro Arg Asn Ser Asp Pro Ala Leu Gly Leu Asp Asp 115 120 125 Asp Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu Leu Arg Ala Arg Ala Leu Leu Leu Arg Ala Pro Ala Leu Leu Arg Ala Arg Ala Leu Leu Arg Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu Leu Arg Ala Arg Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu Leu Arg Ala Arg Ala Pro Ala Ala Gln Leu Ala Arg Ala Leu Leu Arg Ala	

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Gly Pro Asp Ala Glu	Glu Ala Gly A	Asp Glu Thr Pro	Asp Val Asp Pro
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Glu Leu Leu Arg Tyr	Leu Leu Gly A	Arg Ile Leu Ala	Gly Ser Ala Asp
195	200		205
Ser Glu Gly Val Ala	Ala Pro Arg A	Arg Leu Arg Arg	Ala Ala Asp His
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Asp Val Gly Ser Glu	Leu Pro Pro (Glu Gly Val Leu	Gly Ala Leu Leu
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Arg Val Lys Arg Leu	Glu Thr Pro	Ala Pro Gln Val	Pro Ala Arg Arg
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<212> DNA

<213> Homo sapiens

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<400> 102

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<211> 915

<212> DNA

(213) Homo sapiens

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⟨210⟩ 104

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<212> DNA

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(400> 104

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<211> 1323

<212> DNA

<213> Homo sapiens

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⟨210⟩ 106

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<213> Homo sapiens

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<213> Homo sapiens

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<213> Homo sapiens

220/307

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(213) Homo sapiens

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gc	cgcgg	gcgc	tccg	accc	cg g	gcccc	cggt	c ta	cgac	gacg	gcc	ccgc	ggg	ccce	gatg	ct 540
ga	ggagg	cag	gcga	cgag	ac a	eccg	acgt	g ga	cccc	gágc	tgt	tgag	gta	cttg	ctgg	ga 600
cg	gatic	ttg	cggg	aagc	gc g	gact	ccga	g gg	ggtg	gcag	ccc	cgcg	ccg	cctc	cgcc	gt 660
gc	cgcce	gacc	acga	tgtg	gg c	tctg	agct	g cc	ccct	gagg	gcg	tgct	ggg	ggcg	ctgc	tg 720
cg	tgtga	aac	gcct	agag	ac c	ccgg	cgcc	с са	ggtg	cctg	cac	gccg	cct	cttg	ccac	cc 780
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		Met	Val	Gly	Ala	Met	Trp	Lys	Val	Ile	Val	Ser	Leu	Val	Leu	
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Leu	ı Met	Pro	Gly	Pro	Cys	Asp	Gly	Leu	Phe	Arg	Ser	Leu	Tyr	Arg	Ser	
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Va]	Ser	Met	Pro	Pro	Lys	Gly	Asp	Ser	Gly	Gln	Pro	Leu	Phe	Leu	Thr	
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cct	tac	att	gaa	gct	ggg	aag	atc	caa	aaa	gga	aga	gaa	ttg	agt	ttg	253

Pro Tyr Ile Glu A	ala Gly Lys Ile	Gln Lys Gly A	Arg Glu Leu Ser	Leu
50		55	60	
gtc ggc cct ttc c	cca gga ctg aac	atg aag agt (tat gcc ggc ttc	ctc 301
Val Gly Pro Phe I	Pro Gly Leu Asn	Met Lys Ser 1	Tyr Ala Gly Phe	Leu
65	70)	7 5	
acc gtg aat aag	act tac aac ago	aac ctc ttc	ttc tgg ttc ttc	cca 349
Thr Val Asn Lys				
80	85		90	
gct cag ata cag	cca gaa gat gco	c cca gta gtt	ctc tgg cta cag	ggt 397
Ala Gln Ile Gln				
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ggg ccg gga ggt		t gga ctc ttt	gtg gaa cat ggg	cct 445
Gly Pro Gly Gly				
dly (10 dly dl)	115	120	125	
tat gtt gtc aca			aga gac ttc ccc	tgg 493
Tyr Val Val Thr				
	Ser Ash Met In	135	140	•
130				ggc 541
acc aca acg ctc		•		
Thr Thr Thr Leu				Uly
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Phe Ser Phe Thr	Asp Asp Thr Hi	is Gly Tyr Ala	Val Asn Glu Asp	o Asp
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Val Ala Arg Asp	Leu Tyr Ser Al	la Leu Ile Gln	Phe Phe Gln Ile	e Phe

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Pro	Glu	Tyr	Lys	Asn	Asn	Asp	Phe	Tyr	Val	Thr	Gly	Glu	Ser	Tyr	Ala	
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Gly	Lys	Tyr	Val	Pro	Ala	Ile	Ala	His	Leu	lle	His	Ser	Leu	Asn	Pro	
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Val	Arg	Glu	Val	Lys	Ile	Asn	Leu	Asn	Gly	lle	Ala	Ile	Gly	Asp	Gly	
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tat	tct	gat	ccc	gaa	tca	att	ata	ggg	ggc	tat	gca	gaa	ttc	ctg	tac	829
Tyr	Ser	Asp	Pro	Glu	Ser	Ile	Ile	Gly	Gly	Tyr	Ala	Glu	Phe	Leu	Tyr	
	240					245					250					
caa	att	ggc	ttg	ttg	gat	gag	aag	caa	aaa	aag	tac	ttc	cag	aag	cag	877
Gln	Ile	Gly	Leu	Leu	Asp	Glu	Lys	Gln	Lys	Lys	Tyr	Phe	G1n	Lys	Gln	
255					260	١				265	•				270	
tgc	cat	gaa	tgc	ata	gaa	cac	atc	agg	aag	cag	aac	tgg	ttt	gag	gcc	925
Cys	His	Glu	Cys	Ile	Glu	His	Ile	Arg	Lys	Gln	Asn	Trp	Phe	e Glu	Ala	
	•			275	•				280	}				285	i	
ttt	gaa	ata	ctg	gat	aaa	cta	cta	gat	ggc	gac	tta	aca	agt	t gat	cct	973
Phe	Glu	Ile	Leu	Asp	Lys	Leu	Leu	ı Asp	Gly	Asp	Leu	Thr	Ser	r Asp	Pro	
			290)				295	•				300)		
tct	tac	tto	cag	g aat	gtt	aca	gga	tgt	agt	aat	tac	: tat	aac	ttt	ttg	102
Ser	Tyr	Phe	Gln	n Asr	ı Val	Thr	Gly	Cys	Ser	· Asr	1 Tyr	Tyr	- Ası	n Phe	e Leu	
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cgg	tgc	acg	gaa	cct	gag	gat	cag	ctt	tac	tat	gtg	aaa	ttt	ttg	tca	1069
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ctc	cca	gag	gtg	aga	caa	gcc	atc	cac	gtg	ggg	aat	cag	act	ttt	aat	1117
Leu	Pro	Glu	Val	Arg	Gln	Ala	Ile	His	Val	Gly	Asn	Gln	Thr	Phe	Asn	
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gat	gga	act	ata	gtt	gaa	aag	tac	ttg	cga	gaa	gat	aca	gta	cag	tca	1165
Asp	Gly	Thr	Ile	Val	Glu	Lys	Tyr	Leu	Arg	Glu	Asp	Thr	Val	Glr	Ser	
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gtt	aag	cca	tgg	tta	act	gaa	atc	atg	aat	aat	tat	aag	g gt1	t ctg	g atc	1213
Val	Lys	Pro	Trp	Leu	Thr	Glu	Ile	Met	Asn	Asn	Tyr	Lys	s Vai	l Lei	ı Ile	
			370)				375					380	0		
tac	aat	gg(c caa	cte	gac	ato	ato	gtg	gca	gct	gco	ct	gac	a ga	g cac	1261
Tyr	Ası	ı Gl	y Glı	ı Lei	ı Asp	Ile	: I1e	e Val	Ala	Ala	Ala	a Le	u Th	r Gl	u His	;
		38	5				390)				39	5			
															g gca	
Sei	r Le	u Me	t Gl	y Me	t Ası	o Trį	p Ly:	s Gly	y Sei	Glr	n Gl	u Ty	r Ly	s Ly	s Ala	3
	40	0				40	5				41	0				
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Gl	u Ly	s Ly	's Va	l Tr	p Ly	s Il	e Ph	e Ly	s Se	r Ası	p Se	r Gl	u Va	al Al	a Gl	
41					42					42					43	
															gt gg	
Ту	r Il	e Ar	rg Gl	n Al	a Gl	y As	p Ph	e Hi	s Gl	n Va	1 II	.e I	le Ai		ly Gl	У
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ge	ga ca	it a	tt tt	a co	c ta	it ga	ic ca	ag co	t ct	g ag	a go	t t	tt g	ac a	tg at	t 1453

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aat cga ttc att tat gga aaa gga tgg gat cct tat gtt gga taaac	1500
Asn Arg Phe Ile Tyr Gly Lys Gly Trp Asp Pro Tyr Val Gly	
465 470 475	
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ctctatcggc tgcatgccta gaccaccaaa gcgttctgac cggacagtgt cactggagaa	180
ggcggcgcga c atg tcc agg gcg cag atc tgg gct ctg gtg tct ggt gtc	230
Met Ser Arg Ala Gln Ile Trp Ala Leu Val Ser Gly Val	
1 5 10	
gga ggg ttt gga gct ctc gtt gct gct acc acg tcc aat gag tgg aaa	278
Cl., Cl., Pho Cl., Ala Leu Val Ala Ala Thr Thr Ser Asn Glu Trp Lys	

	15					20					25						
gtg	acc	acg	cga	gcc	tcc	tcg	gtg	ata	aca	gcc	act	tgg	gtt	tac	cag	32	26
Val	Thr	Thr	Arg	Ala	Ser	Ser	Val	Ile	Thr	Ala	Thr	Trp	Val	Tyr	Gln		•
30					35					40					45		
ggt	ctg	tgg	atg	aac	tgc	gca	ggt	aac	gcg	ttg	ggt	tct	ttc	cat	tgc	37	74
Gly	Leu	Trp	Met	Asn	Cys	Ala	Gly	Asn	Ala	Leu	Gly	Ser	Phe	His	Cys		
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cga	ccg	cat	ttt	act	atc	ttc	aaa	gta	gca	ggt	tat	ata	cag	gca	tgt	42	22
Arg	Pro	His	Phe	Thr	Ile	Phe	Lys	Val	Ala	Gly	Tyr	Ile	Gln	Ala	Cys		
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aga	gga	ctt	atg	atc	gct	gct	gtc	agc	ctg	ggc	ttc	ttt	ggt	tcc	ata	4	70
Arg	Gly	Leu	Met	Ile	Ala	Ala	Val	Ser	Leu	Gly	Phe	Phe	Gly	Ser	Ile		
		80					85					90					
ttt	gcg	ctc	ttt	gga	atg	aag	tgt	acc	aaa	gtc	gga	ggc	tcc	gat	aaa	5	18
Phe	Ala	Leu	Phe	Gly	Met	Lys	Cys	Thr	Lys	Val	Gly	Ġly	Ser	Asp	Lys		
	95					100					105						
gcc	aaa	gct	aaa	att	gct	tgt	ttg	gct	ggg	att	gta	ttc	ata	ctg	tca	5	66
Ala	Lys	Ala	Lys	Ile	Ala	Cys	Leu	Ala	Gly	Ile	Val	Phe	Ile	Leu	Ser		
110					115				٠	120		•			125		
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Gly	Leu	Cys	Ser	Met	Thr	Gly	Cys	Ser	Leu	Tyr	Ala	Asn	Lys	Ile	Thr		
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acg	gaa	ttc	ttt	gat	cct	ctc	ttt	gtt	gag	caa	aag	tat	gaa	tta	gga	6	62
Thr	Glu	Phe	Phe	Asp	Pro	Leu	Phe	Val	Glu	Gln	Lys	Tyr	Glu	Leu	Gly		
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227/307

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Gly	Val	Ile	Phe	Cys	Phe	Ser	Ile	Ser	Asp	Asn	Asn	Lys	Thr	Pro	Arg	
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Tyr	Thr	Tyr	Asn	Gly	Ala	Thr	Ser	Val	Met	Ser	Ser	Arg	Thr	Lys	Tyr	
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His	Gly	Gly	Glu	ı Asp	Phe	Lys	Thr	Thr	Asn	Pro	Ser	Lys	Gln	Phe	. Asp	
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Lys	. Asr	ı Ala	а Туг	r Val												
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Val Leu Leu Ala Ser Leu Gly Val Gly Leu Val Thr Leu Leu Gly Leu	I
10 15 20	
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Ala Val Gly Ser Tyr Leu Val Arg Arg Ser Arg Arg Pro Gln Val Thr	•
25 30 35	
ctc ctg gac ccc aat gaa aag tac ctg cta cga ctg cta gac aag acg	g 198
Leu Leu Asp Pro Asn Glu Lys Tyr Leu Leu Arg Leu Leu Asp Lys Thr	.
40 45 50 55	5
act gtg agc cac aac acc aag agg ttc cgc ttt gcc ctg ccc acc gcc	246
Thr Val Ser His Asn Thr Lys Arg Phe Arg Phe Ala Leu Pro Thr Ala	3
60 65 70	
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His His Thr Leu Gly Leu Pro Val Gly Lys His Ile Tyr Leu Ser Th	r
75 80 85	
cga att gat ggc agc ctg gtc atc agg cca tac act cct gtc acc ag	t 342
Arg Ile Asp Gly Ser Leu Val Ile Arg Pro Tyr Thr Pro Val Thr Se	r
90 95 100	
gat gag gat caa ggc tat gtg gat ctt gtc atc aag gtc tac ctg aa	g 390

Asp Glu Asp Gln Gly Tyr Val Asp Leu Val Ile Lys Val Tyr Leu Lys	
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ggt gtg cac ccc aaa ttt cct gag gga ggg aag atg tct cag tac ctg	438
Gly Val His Pro Lys Phe Pro Glu Gly Gly Lys Met Ser Gln Tyr Leu	
120 125 130 135	
gat agc ctg aag gtt ggg gat gtg gtg gag ttt cgg ggg cca agc ggg	486
Asp Ser Leu Lys Val Gly Asp Val Val Glu Phe Arg Gly Pro Ser Gly	
140 145 150	
ttg ctc act tac act gga aaa ggg cat ttt aac att cag ccc aac aag	534
Leu Leu Thr Tyr Thr Gly Lys Gly His Phe Asn Ile Gln Pro Asn Lys	
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aaa tot cca cca gaa ccc cga gtg gcg aag aaa ctg gga atg att gcc	582
Lys Ser Pro Pro Glu Pro Arg Val Ala Lys Lys Leu Gly Met Ile Ala	
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Gly Gly Thr Gly Ile Thr Pro Met Leu Gln Leu Ile Arg Ala Ile Leu	
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Lys Val Pro Glu Asp Pro Thr Gln Cys Phe Leu Leu Phe Ala Asn Gln	•
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Arg Tyr Pro Asn Arg Phe Lys Leu Trp Phe Thr Leu Asp His Pro Pro	

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Glu His	Leu	Pro	Ala	Pro	Gly	Asp	Asp	Val	Leu	Val	Leu	Leu	Cys	Gly	
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cca ccc	cca	atg	gtg	cag	ctg	gcc	tgc	cat	ccc	aac	ttg	gac	aaa	ctg	918
Pro Pro	Pro	Met	Val	Gln	Leu	Ala	Cys	His	Pro	Asn	Leu	Asp	Lys	Leu	
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ggc tac	tca	caa	aag	atg	cga	ttc	acc	tac	tg	agca	tcct	cc a	gctt	ccctg	970
Gly Ty	r Ser	Gln	Lys	Met	Arg	Phe	Thr	Tyr					•	٠.	
			300	١				305)						
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	Ме	et Gly Trp Thr Met	
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agg ctg gtc aca gca gca	ctg tta ctg ggt ctc	atg atg gtg gtc act	161
Arg Leu Val Thr Ala Ala	Leu Leu Leu Gly Leu	Met Met Val Val Thr	
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gga gac gag gat gag aad	age eeg tgt gee cat	gag gcc ctc ttg gac	209
Gly Asp Glu Asp Glu Asr	s Ser Pro Cys Ala His	Glu Ala Leu Leu Asp	
25	30	35	
gag gac acc ctc ttt tge	c cag ggc ctt gaa gtt	ttc tac cca gag ttg	257
Glu Asp Thr Leu Phe Cy	s Gln Gly Leu Glu Val	Phe Tyr Pro Glu Leu	
40	45	50	
ggg aac att ggc tgc aa	g gtt gtt cct gat tgt	aac aac tac aga cag	305
Gly Asn Ile Gly Cys Ly	s Val Val Pro Asp Cys	s Asn Asn Tyr Arg Gln	
55	60	65	
aag ate are tee tgg at	g gag ccg ata gtc aag	ttc ccg ggg gcc gtg	353

	Val	Ala	Gly	Pro	Phe	Lys	Val	Ile	Pro	Glu	Met	Trp	Ser	Thr	Ile	Lys
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401	agc	cct	gcc	gat	cca	gat	gtg	atg	gtg	ctg	atc	tat	acc	gca	ggc	gac
	Ser	Pro	Ala	Asp	Pro	Asp	Val	Met	Val	Leu	Ile	Tyr	Thr.	Ala	Gly	Asp
		100					95					90				
449	gat	aca	gta	ctg	tgg	cat	aga	tgg	tťc	aga	cag	aga	ccc	gaa	gca	aga
	Asp	Thr	Val	Leu	Trp	His	Arg	Trp	Phe	Arg	Gln	Arg	Pro	Glu	Ala	Arg
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497	tta	gag	cag	ggc	cag	att	aag	ggg	aaa	aag	ctg	gac	gcc	ggc	aag	atc
	Leu	Glu	Gln	Gly	Gln	Ile	Lys	Gly	Lys	Lys	Leu	Asp	Ala	Gly	Lys	Ile
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545	cgc	cat	ttc	ggc	agt	cac	gca	ccg	cca	tcc	ccc	gct	cag	tac	gcc	tca
	Arg	His	Phe	Gly	Ser	His	Ala	Pro	Pro	Ser	Pro	Ala	Gln	Tyr	Ala	Ser
					145					140				ı	135	
593	ctt	ctc	tct	atc	gtc	aaa	gga	gaa	cag	ctt	tat	gto	ttt	ttc	cag	tac
	Leu	Leu	Ser	Ile	Val	Lys	Gly	Glu	Gln	Leu	Tyr	Val	Phe	Phe	Gln	Tyr
	165					160				i	155)	150
641	ctg	ttt	aga	gac	atg	aaa	tgg	tct	ggo	cga	act	aaa	aac	g gaa	aag	cco
٠	Leu	Phe	Arg	Asp	Meț	Lys	Trp	Ser	Gly	Are	Thr	Lys	Asn	Glu	Lys	Pro
)	180					175)	170			٠	
689	acc	ate	tto	cag	acc	ago	gca	gaa	cct	gaa	ggg	cte	cac	tto	cgt	aad
	. Thr	Met	Phe	Gln	Thr	Ser	Ala	Glu	ı Pro	Glu	ı Gly	: Lei	His	g Phe	n Arg	Ası
		i	195)	190				5	185			
737	g gcc	agg	gaa	aga	ccc	gct	cag	cto	a aco	a cca	c tca	g gao	cag	c tac	g aad	ca
	g Ala	Ar	Glu	Are	Pro	Ala	. Glr	r Lei	. Thi	- Pr	5 Sei	Δer	. G1.	. т.,,		C1.

200	205	210	
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Ser Glu Pro Lys His Lys As	n Gln Ala Glu Ile Ala	Ala Cys	
215 22	225		
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atggaacccc ctctggatac agaa	cccctt cttttccaaa taa	aaaaaaa atcatcc 897	
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ctcccatctc cctcaccagc cgg			
gagcciggcc igggagccag g a	tg gcc atc cac aaa gcc	ttg gtg atg tgc 171	
	et Ala Ile His Lys Ala		
	1 5	10	
ctg gga ctg cct ctc ttc c	tg ttc cca ggg gcc tgg	g gcc cag ggc cat 219	
Leu Gly Leu Pro Leu Phe L			
15	20	25	
gtc cca ccc ggc tgc agc c	aa ggc ctc aac ccc cti	g tac tac aac ctg 267	
Val Pro Pro Gly Cys Ser (
,	•		

			.30					35					40			
tgt :	gac	cgc	tct	ggg	gcg	tgg (ggc :	atc	gtc	ctg	gag	gcic	gtg	gct	ggg	315
					Ala											
		45			·		50					55				
gcg	ggc	att	gtc	acc	acg	ttt	gtg	ctc	acc	atc	atc	ctg	gtg	gcc	agc	363
Ala	Gly	Ile	Val	Thr	Thr	Phe	Val	Leu	Thr	Ile	Ile	Leu	Val	Ala	Ser	
	60					65					70					
ctc	ccc	ttt	gtg	cag	gac	acc	aag	aaa	cgg	agc	ctg	ctg	ggg	acc	cag	411
Leu	Pro	Phe	Val	Gln	Asp	Thr	Lys	Lys	Arg	Ser	Leu	Leu	Gly	Thr	G1n	ı
75					80					85					90)
gta	ttc	ttc	ctt	ctg	ggg	acc	ctg	ggc	ctc	ttc	tgc	cto	gtg	ttt	gco	459
Val	Phe	Phe	Leu	Leu	Gly	Thr	Leu	Gly	Leu	Phe	Cys	Leu	ı Val	Phe	Ala	ı
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tgt	gtg	gtg	g aag	ccc	gac	ttc	tcc	acc	tgt	gcc	tct	. cg	g cgo	c tte	cto	507
Cys	Val	Val	Lys	Pro	Asp	Phe	Ser	Thr	Cys	Ala	Ser	Arı	g Ar	g Pho	e Lei	u
			110)				115	i				120	0		
ttt	gge	g gt	t ctį	g tto	gcc	atc	tgc	tto	tct	tgt:	ct	g gc	g gc	t ca	c gt	с 555
Phe	Gly	/ Val	l Lei	ı Phe	e Ala	Ile	Cys	Phe	Ser	Cys	s Le	u Al	a Al	a Hi	s Va	1
		12	5				130)				13	5			
ttt	gc	cct	c aa	c tte	c cte	g gcc	cgg	g aag	g aac	c ca	c gg	g cc	c cg	g gg	c tg	g 603
Phe	Al:	a Le	u As	n Pho	e Lei	ı Ala	Arg	g Ly:	s Asr	n Hi	s Gl	y Pr	o Ar	g Gl	y Tr	p
	14	0				145	6				15	0				
gte	gat	c tt	c ac	t gt	g gc	t ct	cte	g ct	g ac	c ct	g gt	a ga	g gt	c at	c at	.c 651
Val	11	e Ph	e Th	r Va	1 Al:	a Lei	ı Lev	ı Le	u Th	r Le	u Va	1 G1	u Va	al II	le Il	le .
15	5				16	0				16	5				17	70

699	ggc	gag	ggc	agt	ggc	cgg	gtt	ctg	acc	atc	atc	ctg	tgg	gag	aca	aat
	Gly.	Glu	Gly	Ser	Gly	Arg	Val	Leu	Thr	Ile	Ile	Leu	Trp	Glu	Thr	Asn
		185					180		٠			175				
747	tgt	ccc	tcc	gcc	gtg	gcc	tgg	ggc	gca	agc	agc	aac	ggc	cag	cct	ggc
	Cys	Pro	Ser	Ala	Val	Ala	Trp	Gly	Ala	Ser	Ser	Asn	Gly	G1n	Pro	Gly
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795	ctg	atg	gtc	tac	atc	ctc	gca	atg	gtc	ttt	gac	atg	aac	gcc	atc	gcc
	Leu	Met	Val	Tyr	Ile	Leu	Ala	Met	Val	Phe	Asp	Met	Asn	Ala	Ile	Ala
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843	cgc	ggc	tgt	ctg	gcc	ccc	tgg	gcc	ggg	ctg	ttc	gcc	ggt	ctg	ctg	ctg
	Arg	Gly	Cys	Leu	Ala	Pro	Trp	Ala	Gly	Leu	Phe	Ala	Gly	Leu	Leu	Leu
					230					225					220	
891	gcc	aca	acc	ctc	ctc	gtg	ttt	gtc	ggg	cat	aag	cgt	tgg	cgc	aag	tac
	Ala	Thr	Thr	Leu	Leu	Val	Phe	Val	Gly	His	Lys	Arg	Trp	Arg	Lys	Tyr
	250					245				•	240					235
939	ggc	tac	act	tat	atg	gtc	atc	tgg	gtg	gtg	tgg	ata	gcc	gtt	tcc	acc
	Gly	Tyr	Thr	Tyr	Met	Val	Ile	Trp	Val	Val	Trp	Ile	Ala	Val	Ser	Thr
	1	265					260					255				
987	atc	gcc	ctg	acg	ccc	gac	gat	tgg	acc	ccc	agt	aac	cac	cag	aag	aac
	Ile	Ala	Leu	Thr	Pro	Asp	Asp	Trp	Thr	Pro	Ser	Asn	His	G1n	Lys	Asn
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1035	ccc	ato	gto	tac	ttc	ctc	gtc	tto	gco	tgg	gco	aat	gcc	gcc	cto	gcc
	Pro	Ile	Val	Tyr	Phe	Leu	. Val	Phe	Ala	Trp	Ala	Asn	Ala	Ala	Leu	Ala
			,	295)	290				5	285		
1083	g ggg	cag	tac	ago	caa	gag	cca	ago	z tco	aag	aco	ete	cae	tee	øto	gag

lu	Val	Ser	Gln	Val	Thr	Lys	Ser	Ser	Pro	Glu	Gln	Ser	Tyr	Gln	Gly	
	300					305					310					
gac	atg	tac	ccc	acc	cgg	ggc	gtg	ggc	tat	gag	acc	atc	ctg	aaa	gag	11,31
Asp	Met	Tyr	Pro	Thr	Arg	Gly	Val	Gly	Tyr	Glu	Thr	Ile	Leu	Lys	Glu	
315					320					325	•				330	
cag	aag	ggt	cag	agc	atg	ttc	gtg	gag	aac	aag	gcc	ttt	tcc	atg	gat	1179
Gln	Lys	Gly	Gln	Ser	Met	Phe	Val	Glu	Asn	Lys	Ala	Phe	Ser	Met	Asp	
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gag	ccg	gtt	gca	gct	aag	agg	ccg	gtg	tca	cca	tac	agc	ggg	tac	aat	1227
Glu	Pro	Val	Ala	Ala	Lys	Arg	Pro	Val	Ser	Pro	Tyr	Ser	Gly	Tyr	Asn	
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ggg	cag	ctg	ctg	acc	agt	gtg	tac	cag	ccc	act	gag	atg	gcc	ctg	atg	1275
Gly	Gln	Leu	Leu	Thr	Ser	Val	Tyr	Gln	Pro	Thr	Glu	Met	Ala	Leu	Met	
		365	,				370)				375	,			
cac	aaa	gtt	ccg	tcc	gaa	gga	gct	tac	gac	ato	ato	cto	cca	cgg	gcc	1323
His	Lys	. Val	Pro	Ser	Glu	Gly	Ala	Tyr	Asp	Ile	Ile	Leu	Pro	Arg	, Ala	
	380)				385	•				390)				
acc	gco	aac	ago	cag	gtg	ate	ggc	agt	gcc	aac	tcg:	aco	cte	g cgg	gct	1371
Thr	Ala	a Asr	ı Sei	Gln	Val	Met	Gly	Ser	Ala	. Asr	ı Ser	Thi	. Lei	ı Arg	g Ala	
395	i				400)				405	5				410	
gaa	gao	ate	g tac	tcg:	gco	cag	gago	cac	cag	gce	g gco	aca	a cc	g cci	g aaa	1419
Glu	ı Asp	o Mei	t Tyı	: Ser	Ala	Glr	ı Sei	r His	Glr	n Ala	a Ala	1 Th	r Pr	o Pro	Lys	
				415	5				420)				42	5	
gac	gg	c aa	g aad	e to	ca ₈	ggto	e tt	t aga	a aad	ccc	c ta	c gt	g tg	g ga	C	1464
Asp	G1;	y Ly:	s Ası	n Sei	Gli	n Va	l Ph	e Arg	g Ası	n Pro	о Ту	r Va	l Tr	p As	р	

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430 435 440 tgagtc agcggtggcg aggagggcg gtcggatttg gggagggccc tgaggacctg 1520 gccccgggca agggactete caggeteete etececetgg caggeccage aacatgtgee 1580 ccagatgtgg aagggcctcc ctctctgcca gtgtttgggt gggtgtcatg ggtgtccca 1640 1700 cccactcctc agtgtttgtg gagtcgagga gccaacccca gcctcctgcc aggatcacct cggcggtcac actccagcca aatagtgttc tcggggtggt ggctgggcag cgcctatgtt 1760 1820 tetetggaga tteetgeaac etcaagagae tteecaggeg etcaggeetg gatettgete 1866 ctctgtgagg aacaagggtg cctaataaat acatttctgc tttatt <210> 116 <211> 2198 <212> DNA <213> Homo sapiens ⟨220⟩ <221> CDS ⟨222⟩ (50)... (847) <400> 116 55 aaaatggcgt agagcctagc aacagcgcag gctcccagcc gagtccgtt atg gcc Met Ala 1 103 gct gcc gtc ccg aag agg atg agg ggg cca gca caa gcg aaa ctg ctg Ala Ala Val Pro Lys Arg Met Arg Gly Pro Ala Gln Ala Lys Leu Leu 5 10 15 ccc ggg tcg gcc atc caa gcc ctt gtg ggg ttg gcg cgg ccg ctg gtc 151

Pro Gly Ser Ala Ile Gln Ala Leu Val Gly Leu Ala Arg Pro Leu Val

	20					25					30					
ttg	gcg	ctc	ctg	ctt	gtg	tcc	gcc	gct	cta	tcc	agt	gtt	gta	tca	cgg	. 199
Leu	Ala	Leu	Leu	Leu	Val	Ser	Ala	Ala	Leu	Ser	Ser	Val	Val	Ser	Arg	
35					40					45					50	
act	gat	tca	ccg	agc	cca	acc	gta	ctc	aac	tca	cat	att	tct	acc	cca	247
Thr	Asp	Ser	Pro	Ser	Pro	Thr	Val	Leu	Asn	Ser	His	Ile	Ser	Thr	Pro	
				55					60					65		
aat	gtg	aat	gct	tta	aca	cat	gaa	aac	caa	acc	aaa	cct	tct	att	tcc	295
Asn	Val	Asn	Ala	Leu	Thr	His	Glu	Asn	Gln	Thr	Lys	Pro	Ser	Ile	Ser	
			70					75					80			
caa	atc	agc	acc	acc	ctc	cct	ccc	acg	acg	agt	acc	aag	aaa	agt	gga	343
Gln	Ile	Ser	Thr	Thr	Leu	Pro	Pro	Thr	Thr	Ser	Thr	Lys	Lys	Ser	Gly	
		85					90					95				
gga	gca	tct	gtg	gtc	cct	cat	ccc	tcg	cct	act	cct	ctg	tct	caa	gag	391
Gly	Ala	Ser	Val	Val	Pro	His	Pro	Ser	Pro	Thr	Pro	Leu	Ser	Gln	Glu	
	100					105					110					
gaa	gct	gat	aac	aat	gaa	gat	cct	agt	ata	gag	gag	gag	gat	ctt	ctc	439
Glu	Ala	Asp	Asn	Asn	Glu	Asp	Pro	Ser	Ile	Glu	Glu	Glu	Asp	Leu	Leu	
115					120					125					130	
atg	ctg	aac	agt	tct	cca	tcc	aca	gcc	aaa	gac	act	cta	gac	aat	ggc	487
Met	Leu	Asn	Ser	Ser	Pro	Ser	Thr	Ala	Lys	Asp	Thr	Leu	Asp	Asn	Gly	
				135					140					145		
gat	tat	gga	gaa	cca	gac	tat	gac	tgg	acc	acg	ggc	ccc	agg	gac	gac	535
Asp	Tyr	Gly	Glu	Pro	Asp	Tyr	Asp	Trp	Thr	Thr	Gly	Pro	Arg	Asp	Asp	
			150					155					160			

gac	gag	tct	gat	gac	acc	ttg	gaa	gaa	aac	agg	ggt	tac	atg	gaa	att	583
Asp	Glu	Ser	Asp	Asp	Thr	Leu	Glu	Glu	Asn	Arg	Gly	Tyr	Met.	Glu	Ile	
		165					170					175			•	
gaa	cag	tca	gtg	aaa	tct	ttt	aag	atg	cca	tcc	tca	aat	ata	gaa	gag	631
Glu	Gln	Ser	Val	Lys	Ser	Phe	Lys	Met	Pro	Ser	Ser	Asn	Ile	Glu	Glu	
	180					185					190					
gaa	gac	agc	cat	ttc	ttt	ttt	cat	ctt	att	att	ttt	gct	ttt	tgc	att	679
Glu	Asp	Ser	His	Phe	Phe	Phe	His	Leu	Ile	Ile	Phe	Ala	Phe	Cys	Ile	
195					200					205					210	
gct	gtt	gtt	tac	att	aca	tat	cac	aac	aaa	agg	aag	att	ttt	ctt	ctg	727
Ala	Val	Val	Tyr	Ile	Thr	Tyr	His	Asn	Lys	Arg	Lys	Ile	Phe	Leu	Leu	
				215	•				220	ı				225		
gtt	caa	agc	agg	aaa	tgg	cgt	gat	ggc	ctt	tgt	tcc	aaa	aca	gtg	gaa	775
Val	Gln	Ser	Arg	Lys	Trp	Arg	, Asp	Gly	Leu	Cys	Ser	Lys	Thr	Val	Glu	
			230)				235	5				240)		
tac	cat	cgc	cta	gat	cag	aat	gtt	aat	gag	g gca	ate	cct	tct	tte	g aag	823
Tyr	His	Arg	, Leu	ı Asp	Gln	Asr	ı Val	l Asr	ı Glu	ı Ala	. Met	Pro	Ser	Let	ı Lys	
		245	5				250)				255	5			
att	aco	aat	gat	tat	att	tti	t taa	aagc	acte	gtgat	ttt g	gaati	ttgct	tt		870
Ιle	e Thi	r Asr	ı Asp	Ty1	· Ile	Phe	9									
	260)				268	5									
at	gtaai	tttt	atti	tgcti	tga d	ttt	ttata	at g	atati	tgtg	c aaa	atgt	ttgc	cat	aggcaat	930
tg	gtaci	ttaa	atga	agagı	gtg a	agtc	tctc	tt t	tgcc	ttgg	t gc	tttg	gaaa	tta	aatgtca	990
ca	aacga	agta	tata	aatt	ttt 1	tatc	tgta	ct t	ttag	agct	g ag	ttta	atca	ggt	gtocaaa	a 1050
ati	gtga	gtta	aaca	atta	cct	tata	ttta	са с	tgtt	agtt	t tt	attg	tttt	aga	tttatta	a 1110

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tetgacatat etgttaetge tgactcacat teatteteeg ecattcaaat actattttt	1230
atccacattt ttttttgttc ccaaactgta atgtacaagg atatgtgtga taatgctttg	1290
gatttgagta atatttttt ttcttccaag aaaactgctt tggatatttt tagataattt	1350
aaacataatt taggataatg atattgctca atctgaccac aattttaggt aaaacattaa	1410
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aatatgtaaa tatgtgattt gaaccatggt tgacttacaa gtgtcactac agctttttag	1950
aaaacatagc cctaatatat gttaagcagg acccgggtga gccagtgggc ttgcgcttta	2010
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ccgcc	gct	atg	ctg	ggg	ctg	ctg	gtg	gcg	ttg	ctg	gcc	ctg	ggg	ctc	gct	110
		Met	Leu	Gly	Leu	Leu	Val	Ala	Leu	Leu	Ala	Leu	Gly	Leu	Ala	
		1				5					10	•				
gtc t	tt	gcg	ctg	ctg	gac	gtc	tgg	tac	ctg	gtg	cgc	ctt	ccg	tgc	gcc	158
Val F	Phe	Ala	Leu	Leu	Asp	Val	Trp	Tyr	Leu	Val	Arg	Leu	Pro	Cys	Ala	
15					20					25					30	•
gtg	ctg	cgc	gcg	cgc	ctg	ctg	cag	ccg	cgc	gtc	cgt	gac	ctg	cta	gct	206
Val I	Leu	Arg	Ala	Arg	Leu	Leu	Gln	Pro	Arg	Val	Arg	Asp	Leu	Leu	Ala	
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gag	cag	cgc	ttc	ccg	ggc	cgc	gtg	ctg	ccc	tcg	gac	ttg	gac	ctg	ctg	254
Glu	Gln	Arg	Phe	Pro	Gly	Arg	Val	Leu	Pro	Ser	Asp	Leu	Asp	Leu	Leu	
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ttg	cac	atg	aac	aac	gcg	cgc	tac	ctg	cgc	gag	gcc	gac	ttt	gcg	cgc	302
Leu	His	Met	Asn	Asn	Ala	Arg	Tyr	Leu	Arg	Glu	Ala	Asp	Phe	Ala	Arg	
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gtc	gcg	cac	ctg	acc	cgc	tgc	ggg	gtg	ctc	ggg	gcg	ctg	agg	gag	ttg	350
Val	Ala	His	Leu	Thr	Arg	Cys	Gly	Val	Leu	Gly	Ala	Leu	Arg	Glu	Leu	
	80	ı				85					90					
cgg	gcg	cac	acg	gte	ctg	gcg	gco	tcg	g tgc	gcg	cgc	cac	cgc	cgc	tcg	398
Arg	Ala	His	Thr	Val	Leu	ı Ala	Ala	s Ser	Cys	Ala	Arg	His	Arg	Arg	Ser	
95					100)				105	i				110	

ctg	cgc	ctg	ctg	gag	ccc	ttc	gag	gtg	cgc	acc	cgc	ctg	ctg	ggc	tgg	446
Leu	Arg	Leu	Leu	Glu	Pro	Phe	Glu	Val	Arg	Thr	Arg	Leu	Leu	Gly	Trp	
				115			•		120					125		
gac	gac	cgc	gcg	ttc	tac	ctg	gag	gcg	cgc	ttt	gtc	agc	ctg	cgg	gac	494
Asp	Asp	Arg	Ala	Phe	Tyr	Leu	Glu	Ala	Arg	Phe	Val	Ser	Leu	Arg	Asp	
			130					135					140			
ggt	ttc	gtg	tgc	gcg	ctg	ctg	cgc	ttc	ċgg	cag	cac	ctg	ctg	ggc	acc	542
Gly	Phe	Val	Cys	Ala	Leu	Leu	Arg	Phe	Arg	Gln	His	Leu	Leu	G1y	Thr	
		145					150					155				
tca	ccc	gag	cgc	gtc	gtg	cag	cac	ctg	tgc	cag	cgc	agg	gtg	gag	ccc	590
Ser	Pro	Glu	Arg	Val	Val	Gln	His	Leu	Cys	Gln	Arg	Arg	Val	Glu	Pro	
	160					165	•				170)				
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Pro	Glu	Leu	Pro	Ala	Asp	Leu	ı Gln	His	Trp	Ile	Ser	Tyr	- Asr	ı Glu	ı Ala	
175					180)				185	,				190	
agc	ago	cag	ctg	cto	cgc	ate	g gag	g agt	ggg	cto	agt	; gai	t gto	caco	aag	686
Ser	Ser	Gln	Leu	ı Lev	ı Arg	g Mei	t Glu	ı Sei	Gly	Leu	ı Ser	. Ası	y Va	l Thi	, Lys	
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Asp	Glr	1														
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											M	let A	la T	rp A	rg	
		,										1				
cgg	cgc	gaa	gcc	agc	gtc	ggg	gct	cgc	ggc	gtg	ttg	gct	ctg	gcg	ttg	162
Arg	Arg	Glu	Ala	Ser	Val	Gly	Ala	Arg	Gly	Val	Leu	Ala	Leu	Ala	Leu	
5					10		٠			15					20	
ctc	gcc	ctg	gcc	ctg	tgc	gtg	ccc	ggg	gcc	cgg	ggc	cgg	gct	ctc	gag	210
Leu	Ala	Leu	Ala	Leu	Cys	Val	Pro	Gly	Ala	Arg	Gly	Arg	Ala	Leu	Glu	
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tgg	ttc	tcg	gcc	gtg	gta	aac	atc	gag	tac	gtg	gac	ccg	cag	acc	aac	258
Trp	Phe	Ser	Ala	Val	Val	Asn	Ile	Glu	Tyr	Val	Asp	Pro	Gln	Thr	Asn	
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Leu	Thr	Val	Trp	Ser	Val	Ser	Glu	Ser	Gly	Arg	Phe	Gly	Asp	Ser	Ser	
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ccc	aag	gag	ggc	gcg	cat	ggc	ctg	gtg	ggc	gtc	ccg	tgg	gcg	ccc	ggc	354
Pro	Lys	Glu	Gly	Ala	His	Gly	Leu	Val	Gly	Val	Pro	Trp	Ala	Pro	Gly	
	70	i				75					80)				
gga	gac	cto	gag	ggc	tgc	gcg	ccc	gac	acg	cgc	ttc	ttc	gtg	ccc	gag	402
Gly	Asp	Lei	Glu	Gly	Cys	Ala	Pro	Asp	Thr	Arg	Phe	Phe	. Val	Pro	Glu	
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Pro	Gly	Gly	Arg	Gly	Ala	Ala	Pro	Trp	Val	Ala	Leu	Val	Ala	Arg	Gly	
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Gly	Cys	Thr	Phe	Lys	Asp	Lys	Val	Leu	Val	Ala	Ala	Arg	Arg	Asn	Ala	
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Ser	Ala	Val	Val	Leu	Tyr	Asn	Glu	Glu	Arg	Tyr	Gly	Asn	Ile	Thr	Leu	
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ccc	ate	tct	cac	gcg	gga	aca	gga	aat	ata	gtg	gtc	att	atg	att	agc	594
Pro	Met	. Sei	His	s Ala	Gly	Thr	Gly	Asn	Ile	Val	Val	Ile	Met	Ile	Ser	
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Tyr	Pro	Ly:	s Gl	y Arg	g Glu	ı Ile	e Leu	Glu	Leu	Val	G1n	Lys	Gly	, Ile	e Pro	
165					170)				175	5				180	
gta	ac	g at	g ac	c ata	a gg	g gti	t ggc	acc	cgg	cat	gta	a cag	g ga	g tt	c atc	690
Val	Th	r Me	t Th	r Ile	e Gl	y Vai	l Gly	Thr	Arg	His	s Val	l Glr	n Gl	u Ph	e Ile	
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Ser	- G1	y Gl	n Se	r Va	l Va	1 Ph	e Val	l Ala	Ile	e Ala	a Pho	e Il	e Th	r Me	t Met	
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ati	t at	c to	g tt	a gc	c tg	g ct	a ata	a tti	t tac	ta	t ata	a ca	g cg	t tt	c cta	786
Ιl	e Il	e Se	r Le	eu Al	a Tr	p Le	u Il	e Pho	е Туз	r Ty	r Il	e Gl	n Ar	g Ph	e Leu	
		21	5				22	0				22	5			
		+	+.	st ca	σ at	t oo	a ag	t ca	g ag	c ca	t ag	a aa	a ga	a ac	t aag	834

Tyr	Thr	Gly	Ser	Gln	Ile	Gly	Ser	Gln	Ser	His	Arg	Lys	Glu	Thr	Lys	
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Lys	Val	Ile	Gly	Gln	Leu	Leu	Leu	His	Thr	Vaİ	Lys	His	Gly	Glu	Lys	
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Gly	Ile	Asp	Val	Asp	Ala	Glu	Asn	Cys	Ala	Val	Cys	Ile	Glu	Asn	Phe	
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aaa	gta	aag	gat	att	att	aga	att	ctg	cca	tgc	aag	cat	att	ttt	cat	978
Lys	Val	Lys	Asp	Ile	Ile	Arg	Ile	Leu	Pro	Cys	Lys	His	Ile	Phe	His	
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aga	ata	tgo	att	gac	cca	tgg	ctt	ttg	gat	cac	cga	aca	tgt	cca	atg	1026
Arg	Πe	e Cys	: Ile	. Asp	Pro	Trp	Leu	Leu	. Asp	His	Arg	Thr	Cys	Pro	Met	
		298	5				300)				305	5			•
tgt	aaa	a cti	t gat	gto	ato	aaa	gcc	cta	gga	tat	tgg	g gga	a gag	cct	ggg	1074
Cys	Ly:	s Lei	ı Asp	Val	Ιlε	e Lys	Ala	Let	ı Gly	Tyr	Trp	G13	/ Glu	ı Pro	Gly	
	310	0				315	j				320)				
gat	gt	a ca	g gag	gate	g cct	t gct	. cca	a gaa	a tct	cct	cc1	t gga	a agg	g ga	t cca	1122
Asp	Va.	l Gl	n Glu	u Met	t Pro	Ala	Pro	Glu	ı Sei	Pro	Pro	o Gl	y Ar	g As	p Pro	
325	5				330)				335	5				340	
gc	t gc	a aa	t tt	g ag	t cta	a gct	t tta	a cc	a ga	t gat	t ga	c gg	a ag	t ga	t gag	1170
Ala	a Al	a As	n Lei	u Se	r Lei	u Ala	a Le	u Pr	o Ası	a Ası	p As	p Gl	y Se	r As	p Glu	
				34	5				350	0				35	5	
ag	c ag	t cc	а сс	a tc	a gc	c tc	c cc	t gc	t ga	a tc	t ga	g cc	а са	g tg	t gat	1218
ç.	- 50	r D-	~ D~	~ Sa	- A1	a Se	r Pr	o Al	a Gl	u Se	r Gl	u Pr	o Gl	n Cy	s Asp	

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ggc agg agt gac	tct cgg cat gga	gga ccc atc tcc	tagcacac 1310
Gly Arg Ser Asp S	Ser Arg His Gly	Gly Pro Ile Ser	
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cage atg ttt tge	cca ctg aaa ctc	atc ctg ctg cca	gtg tta ctg gat 169
Met Phe Cys	Pro Leu Lys Leu	Ile Leu Leu Pro	Val Leu Leu Asp
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ſyr	Ser	Leu	Gly	Leu	Asn	Asp	Leu	Asn	Val	Ser	Pro	Pro	Glu	Leu	Thr	
				20					25		•			30	•	
gtc	cat	gtg	ggt	gat	tca	gct	ctg	atg	gga	tgt	gtt	ttc	cag	agc	aca	265
Val	His	Val	Gly	Asp	Ser	Ala	Leu	Met	Gly	Cys	Val	Phe	Gln	Ser	Thr	•
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gaa	gac	aaa	tgt	ata	ttc	aag	ata	gac	tgg	act	ctg	tca	cca	gga	gag	313
Glu	Asp	Lys	Cys	Ile	Phe	Lys	Ile	Asp	Trp	Thr	Leu	Ser	Pro	Gly	Glu	1
		50					55					60)			
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His	Ala	Lys	Asp	Glu	Tyr	Val	Leu	Tyr	Tyr	Tyr	Ser	Asn	Leu	. Sei	r Val	1
	65	,				70					75					
cct	att	gge	g cgo	tto	cag	aac	cgc	gta	cac	ttg	atg	gge	g ga	aa	c tt	a 409
Pro	Ile	Gly	/ Arg	g Phe	e Gln	Asn	Arg	, Val	His	Leu	Met	Gly	/ As	Ası	n Le	u
80)				85	;				90)				9	5
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Cys	. Ası	n Ası	p G 1	y Sei	r Leu	ı Lev	ı Lei	ı Glr	ı Asp	Val	Glr	ı Gl	u Al	a As	p Gl	n
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Gl	/ Th	r Ty	r Il	е Су	s Glu	ı Ile	e Ar	g Lei	ı Ly:	s Gly	y Gli	u Se	r Gl	n Va	l Ph	ıe
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Ly	s Ly	s Al	a Va	l Va	l Le	u Hi	s Va	l Le	u Pr	o Gl	u Gl	u Pr	o Ly	s G	lu Le	eu
		13	0				13	5				14	0			
at	e et	с са	t gt	g gg	t gg	a tt	g at	t ca	g at	g gg	a tg	t gt	t ti	c ca	ag a	gc 601

Met	Val	His	Val	Gly	Gly	Leu	Ile	Gln	Met	Gly	Cys	Val	Phe	Gln	Ser	
	145					150					155				•	
aca	gaa	gtg	aaa	cac	gtg	acċ	aag	gta	gaa	tgg	ata	ttt	tca	gga	cgg	649
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Gly Ser Pro Leu Leu Trp Gly P	ro Arg Ala Gl	y Gly Val Gly	Leu Leu
5	10	15	
gtg ctg ctg ctc ctc ggc ctg t			
Val Leu Leu Leu Gly Leu P	he Arg Pro Pr	o Pro Ala Leu	Cys Ala
20 25		30	
cgg ccg gta aag gag ccc cgc g			
Arg Pro Val Lys Glu Pro Arg G	ly Leu Ser Al	a Ala Ser Pro	Pro Leu

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Ala	Glu	Thr	Gly	Ala	Pro	Arg	Arg	Phe	Arg	Arg	Ser	Val	Pro	Arg	Gly	
				55					60					65		
gag	gcg	gcg	ggg	gcg	gtg	cag	gag	ctg	gcg	cgg	gcg	ctg	gcg	cat	ctg	295
Glu	Ala	Ala	Gly	Ala	Val	Gln	Glu	Leu	Ala	Arg	Ala	Leu	Ala	His	Leu	
			70					75					80			
ctg	gag	gcc	gaa	cgt	cag	gag	cgg	gcg	cgg	gcc	gag	gcg	cag	gag	gct	343
Leu	Glu	Ala	Glu	Arg	Gln	Glu	Arg	Ala	Arg	Ala	Glu	Ala	Gln	Glu	Ala	
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gag	gat	cag	cag	gcg	cgc	gtc	ctg	gcg	cag	ctg	ctg	cgc	gtc	tgg	ggc	391
Glu	Asp	Gln	Gln	Ala	Arg	Val	Leu	Ala	Gln	Leu	Leu	Arg	·Val	Trp	Gly	
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gcc	ccc	cgc	aac	tct	gat	ccg	gct	ctg	ggc	ctg	gac	gac	gad	ccc	gac	439
Ala	Pro	Arg	Asn	Ser	Asp	Pro	Ala	Leu	Gly	Leu	Asp	Asp	Asp	Pro	Asp	
115					120					125					130	
gcg	cct	gca	gcg	cag	ctc	gct	cgc	gct	ctg	ctc	cgc	gcc	cgo	ctt	gac	487
Ala	Pro	Ala	Ala	Gln	Leu	Ala	Arg	Ala	Leu	Leu	Arg	Ala	Are	g Leu	ı Asp	
				135	,		٠		140					145	5	
cct	gcc	gcc	ctc	gca	gcc	cag	ctt	gtc	ccc	gcg	ccc	gto	cco	g gcc	gcg	535
Pro	Ala	Ala	Leu	Ala	Ala	Gln	Leu	ı Val	Pro	Ala	Pro	Val	Pro	o Ala	a Ala	
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Ala	Leu	ı Arg	Pro	Arg	g Pro	Pro	Val	Tyr	Asp	Asp	Gly	y Pro	Al:	a Gly	y Pro	
		165	;				170)				179	5			

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			Leu														
195					200					205					21		
ggg	gtg	gca	gcc	ccg	cgc	cgc	ctc	cgc	cgt	gcc	gcc	gac	cac	gat	gt	g	727
			Ala														
				215	,				220)				225	5		
ggc	tct	gag	ctg	ccc	cct	gag	ggc	gtg	ctg	ggg	gcg	ctg	ctg	g cgt	t gt	g	775
Gly	Ser	Gli	Leu	Pro	Pro	Glu	Gly	/ Val	Leu	Gly	Ala	Lei	ı Lev	ı Arg	g Va	a 1	
			230)				235	,				240)			
aaa	cgo	cta	a gag	gaco	cce	g gcg	cc	c cag	gte	g cct	gca	, cgo	cgc	c ct	c ti	tg	823
Lys	s Arı	g Lei	u Glu	ı Thi	r Pro	o Ala	a Pro	o Glr	ı Val	l Pro	Ala	a Arg	g Arı	g Lei	u L	eu	
		24	5				25	0				25	5				
cca	а сс	ct	gagca	actg	cc c	ggat	cccg	t gc	accc	tggg	acc	caga	agt	gccc	ccg	cca	880
Pro	o Pr	0															
	26																
			agg														940
сс	ctct	cacc	cga	ggat	ccc	tacc	ccct	gg c	ccca	caat	a aa	catg	atct	. gaa	igca	gc	998
<2	10>	121															
<2	11>	337															
<2	12>	PRT															
<2	213>	Homo	sap	iens	5												

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Met	Thr	Ala	Gly	Gly	Gln	Ala	Glu	Ala	Glu	Gly	Ala	.Gly	Gļy	Glu	Pro
1				5					10					15	
Gly	Ala	Ala	Arg	Leu	Pro	Ser	Arg	Val	Ala	Arg	Leu	Leu	Ser	Ala	Leu
			20					25					30		
Phe	Tyr	Gly	Thr	Cys	Ser	Phe	Leu	Ile	Val	Leu	Val	Asn	Lys	Ala	Leu
		35					40					45			
Leu	Thr	Thr	Tyr	Gly	Phe	Pro	Ser	Pro	Ile	Phe	Leu	Gly	Ile	Gly	Gln
	50					55					60				
Met	Ala	Ala	Thr	Ile	Met	Ile	Leu	Tyr	Val	Ser	Lys	Leu	Asn	Lys	Ile
65					70					75					80
Ile	His	Phe	Pro	Asp	Phe	Asp	Lys	Lys	Ile	Pro	Val	Lys	Leu	Phe	Pro
		٠		85					90					95	
Leu	Pro	Leu	Leu	Tyr	Val	Gly	Asn	His	Ile	Ser	Gly	Leu	Ser	Ser	Thr
			100)				105					110	1	
Ser	Lys	Leu	Ser	Leu	Pro	Met	Phe	Thr	Val	Leu	Arg	Lys	Phe	Thr	Ile
		115	;				120)				125	5		
Pro	Leu	Thr	Leu	ı Leu	ı Leu	Glu	Thr	· Ile	Ile	e Leu	Gly	Lys	Gln	Tyr	Ser
	130) .				135	i				140)			
Leu	Asn	ı Ile	lle	e Leu	ı Ser	· Val	Phe	e Ala	Ile	e Ile	e Leu	ı Gly	, Ala	Phe	lle
145	,				150)				155	5				160
Ala	Ala	Gly	Sei	r Asp	Leu	ı Ala	n Phe	e Asr	ı Lei	ı Glu	ı Gl	у Туі	r Ile	e Phe	e Val
				165	5				170)				178	5
Phe	. Leu	ı Asr	ı Ası	o Ile	e Phe	Th:	r Ala	a Ala	a Asr	n Gly	y Va	1 Ty:	r Thi	r Lys	s Gln
			180	0				189	5				190)	

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Lys	Met	Asp	Pro	Lys	Glu	Leu	Gly	Lys	Tyr	Gly	Val	Leu	Phe	Tyr	Ası
		195					200					205			
Ala	Cys	Phe	Met	Ile	Ile	Pro	Thr	Leu	Ile	Ile	Ser	Val	Ser	Thr	Gl
	210					215					220				
Asp	Leu	Gln	Gln	Ala	Thr	Glu	Phe	Asn	Gln	Trp	Lys	Asn	Val	Val	Phe
225					230					235					240
Ile	Leu	Gln	Phe	Leu	Leu	Ser	Cys	Phe	Leu	Gly	Phe	Leu	Leu	Met	Туз
				245					250					255	
Ser	Thr	Val	Leu	Cys	Ser	Tyr	Tyr	Asn	Ser	Ala	Leu	Thr	Thr	Ala	Va:
			260	•	•			265					270		
Val	Gly	Ala	Ile	Lys	Asn	Val	Ser	Val	Ala	Tyr	Ile	Gly	Ile	Leu	H
		275					280					285			
Gly	Gly	Asp	Tyr	Ile	Phe	Ser	Leu	Leu	Asn	Phe	Val	Gly	Leu	Asn	Ile
	290					295					300				
Cys	Met	Ala	Gly	Gly	Leu	Arg	Tyr	Ser	Phe	Leu	Thr	Leu	Ser	Ser	Glı
305					310					315					320
Leu	Lys	Pro	Lys	Pro	Val	Gly	Glu	Glu	Asn	Ile	Cys	Leu	Asp	Leu	Lys
				325					330					335	
Ser															

⟨210⟩ 122

<211> 236

<212> PRT

<213> Homo sapiens

<400	> 12	22													
Met	Ala	Glu	Ala	Glu	Glụ	Ser	Pro	Gly	Asp	Pro	Gly	Thr	Ala	Ser	Pro
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Arg	Pro	Leu	Phe	Ala	Gly	Leu	Ser	Asp	Ile	Ser	Ile	Ser	G1n	Asp	Ile
			20					25					30		
Pro	Val	Glu	Gly	Glu	Ile	Thr	Ile	Pro	Met	Arg	Ser	Arg	Ile	Arg	Glu
		35					40					45			
Phe	Asp	Ser	Ser	Thr	Leu	Asn	Glu	Ser	Val	Arg	Asn	Thr	Ile	Met	Arg
	50					55					60				
Asp	Leu	Lys	Ala	Val	Gly	Lys	Lys	Phe	Met	His	Val	Leu	Tyr	Pro	Arg
65					70					75			•		80
Lys	Ser	Asn	Thr	Leu	Leu	Arg	Asp	Trp	Asp	Leu	Trp	Gly	Pro	Leu	Ile
				85					90					95	
Leu	Cys	Val	Thr	Leu	Ala	Leu	Met	Leu	Gln	Arg	Asp	Ser	Ala	Asp	Ser
			100					105					110		
Glu	Lys	Asp	Gly	Gly	Pro	Gln	Phe	Ala	Glu	Val	Phe	Val	Ile	Val	Trp
		115					120					125			
Phe	Gly	Ala	Val	Thr	Ile	Thr	Leu	Asn	Ser	Lys	Leu	Leu	Gly	Gly	Asn
	130					135		•			140				
Ile	Ser	Phe	Phe	Gln	Ser	Leu	Cys	Val	Leu	Gly	Tyr	Cys	Ile	Leu	Pro
145					150					155					160
Leu	Thr	Val	Ala	Met	Leu	Ile	Cys	Arg	Leu	Val	Leu	Leu	Ala		Pro
				165					170					175	
Gly	Pro	Val	Asn	Phe	Met	Val	Arg	Leu	Phe	Val	Val	Ile	Val	Met	Phe
			180					185					190		

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Ala Trp Ser Ile Val Ala Ser Thr Ala Phe Leu Ala Asp Ser Gln Pro Pro Asn Arg Arg Ala Leu Ala Val Tyr Pro Val Phe Leu Phe Tyr Phe Val Ile Ser Trp Met Ile Leu Thr Phe Thr Pro Gln <210> 123 <211> 560 <212> PRT <213> Homo sapiens <400> 123 Met Ala Ala Pro Ala Glu Ser Leu Arg Arg Arg Lys Thr Gly Tyr Ser Asp Pro Glu Pro Glu Ser Pro Pro Ala Pro Gly Arg Gly Pro Ala Gly Ser Pro Ala His Leu His Thr Gly Thr Phe Trp Leu Thr Arg Ile Val Leu Leu Lys Ala Leu Ala Phe Val Tyr Phe Val Ala Phe Leu Val Ala Phe His Gln Asn Lys Gln Leu Ile Gly Asp Arg Gly Leu Leu Pro Cys Arg Val Phe Leu Lys Asn Phe Gln Gln Tyr Phe Gln Asp Arg Thr Ser Trp Glu Val Phe Ser Tyr Met Pro Thr Ile Leu Trp Leu Met Asp Trp

			100					105					110		
Ser	Asp	Met	Asn	Ser	Asn	Leu	Asp	Leu	Leu	Ala	Leu	Leu	Gly	Leu	Gly
		115		•			120					125			
Ile	Ser	Ser	Phe	Val	Leu	Ile	Thr	Gly	Cys	Ala	Asn	Met	Leu	Leu	Met
	130					135					140				
Ala	Ala	Leu	Trp	Gly	Leu	Tyr	Met	Ser	Leu	Val	Asn	Val	Gly	His	Val
145					150					155					160
Trp	Tyr	Ser	Phe	Gly	Trp	Glu	Ser	G1n	Leu	Leu	Glu	Thr	Gly	Phe	Leu
				165					170					175	
Gly	Ile	Phe	Leu	Cys	Pro	Leu	Trp	Thr	Leu	Ser	Arg	Leu	Pro	Gln	His
			180					185					190		
Thr	Pro	Thr	Ser	۸rg	Ile	Val	Leu	Trp	Gly	Phe	Arg	Trp	Leu	Ile	Phe
		195					200					205			
Arg	Ile	Met	Leu	Gly	Ala	Gly	Leu	Ile	Lys	Ile	Arg	Gly	Λsp	Arg	Cys
	210					215					220				
Trp	Arg	Asp	Leu	Thr	Cys	Met	Asp	Phe	His	Tyr	Glu	Thr	Gln	Pro	
225					230)				235	ı				240
Pro	Asn	Pro	Val	Ala	Tyr	Tyr	Leu	His	His	Ser	Pro	Trp	Trp		
				245					250					255	
Arg	Phe	Glu	Thr	Leu	Ser	Asn	His	Phe	Ile	Glu	Leu	Leu	\Val	Pro	Phe
			260)				265	•				270)	
Phe	Leu	Phe	Leu	Gly	Arg	Arg	Ala	Cys	Ile	Ile	His	Gly	Val	Leu	Glr
		275					280					285			
Ile	Leu	. Phe	Gln	ı Ala	Val	Leu	Ile	Val	Ser	Gly	Asr	ı Lei	ı Ser	Phe	e Leu
	290)				295	•				300)			

Asn	Trp	Leu	Thr	Met	Val	Pro	Ser	Leu	Ala	Cys	Phe	Asp	Asp	Ala	Thr
305					310					315					320
Leu	Gly	Phe	Leu	Phe	Pro	Ser	Gly	Pro	Gly	Ser	Leu	Lys	Asp	Arg	Val
				325			•		330					335	
Leu	Gln	Met	Gln	Arg	Asp	Ile	Arg	Gly	Ala	Arg	Pro	Glu	Pro	Arg	Phe
			340					345					350		
Gly	Ser	Val	Val	Arg	Arg	Ala	Ala	Asn	Val	Ser	Leu	Gly	Val	Leu	Leu
		355					360					365			
Ala	Trp	Leu	Ser	Val	Pro	Val	Val	Leu	Asn	Leu	Leu	Ser	Ser	Arg	Gln
	370					375					380				
Val	Met	Asn	Thr	His	Phe	Asn	Ser	Leu	His	Ile	Val	Asn	Thr	Tyr	Gly
385	·				390					395					400
Ala	Phe	Gly	Ser	Ile	Thr	Lys	Glu	Arg	Ala	Glu	۷aḷ	Ile	Leu	Gln	Gly
				405					410					415	
Thr	Ala	Ser	Ser	Asn	Ala	Ser	Ala	Pro	Asp	Ala	Met	Trp	Glu	Asp	Tyr
			420					425					430		
Glu	Phe	Lys	Cys	Lys	Pro	Gly	Asp	Pro	Ser	Arg	Arg	Pro	Cys	Leu	Ile
		435					440					445			
Ser	Pro	Tyr	His	Tyr	Arg	Leu	Asp	Trp	Leu	Met	Trp	Phe	Ala	Ala _.	Phe
	450					455					460				
Gln	Thr	Tyr	Glu	His	Asn	Asp	Trp	Ile	Ile	His	Leu	Ala	Gly	Lys	Leu
165					470					475					480
_eu	Ala	Ser	Asp	Ala	Glu	Ala	Leu	Ser	Leu	Leu	Ala	His	Asn	Pro	Phe
				485					490					495	
Ma	Glv	Aro	Pro	Pro	Pro	Arø	Trn	Val	Ara	C1v	Glu	Hic	Tvr	Aro	Tvr

			500					505					510		
Lys	Phe	Ser	Arg	Pro	G _l y	Gly	Arg	His	Ala	Ala	Glu	Gly	Lys	Trp	Trp
		515					520					525			
Val	Arg	Lys	Arg	Ile	Gly	Ala	Tyr	Phe	Pro	Pro	Leu	Ser	Leu	Glu	Glu
	530					535					540				
Leu	Arg	Pro	Tyr	Phe	Arg	Asp	Arg	Gly	Trp	Pro	Leu	Pro	Gly	Pro	Leu
545					550					555					560
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<21	1> 4	06													
<21	2> P	RT													
<21	3> H	omo	sapi	ens											
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Met	Ala	Glu	Asn	Gly	Lys	Asn	Cys	Asp	Gln	Arg	Arg	Val	Ala	Met	Asn
1				5					10					15	
Lys	Glu	His	His	Asn	Gly	Asn	Phe	Thr	Asp	Pro	Ser	Ser	Val	Asn	Glu
			20					25					30		
Lys	Lys	Arg	Arg	Glu	Arg	Glu	Glu	Arg	Gln	Asn	Ile	Val	Leu	Trp	Arg
		35					40	•				45	ı		
Gln	Pro	Leu	Ile	Thr	Leu	Gln	Tyr	Phe	Ser	Leu	Glu	Ile	Leu	Val	Ile
	50)				55	,				60)			
Leu	Lys	Glu	Trp	Thr	Ser	Lys	Leu	Trp	His	Arg	Glr	Ser	Ile	Val	Val
65	i				70)				75	,				80
Ser	Phe	Leu	Leu	Leu	ı Lev	Ala	Val	Leu	ı Ile	e Ala	The	Tyr	Tyr	Val	Glu
				85	j				90)				95	i

Gly	Val	His	Gln	Gln	Tyr	Val	Gln	Arg	Ile	Glu	Lys	Gln	Phe	Leu	Leu
			100					105					110		
Tyr	Ala	Tyr	Trp	Ile	Gly	Leu	Gly	Ile	Leu	Ser	Ser	Val	Gly	Leu	Gly
		115					120					125			
Thr	Gly	Leu	His	Thr	Phe	Leu	Leu	Tyr	Leu	Gly	Pro	His	Ile	Ala	Ser
	130					135					140				
Val	Thr	Leu	Ala	Ala	Tyr	Glu	Cys	Asn	Ser	Val	Asn	Phe	Pro	Glu	Pro
145					150					155					160
Pro	Tyr	Pro	Asp	Gln	Ile	Ile	Cys	Pro	Asp	Glu	Glu	Gly	Thr	Glu	Gly
				165					170					175	
Thr	Ile	Ser	Leu	Trp	Ser	Ile	Ile	Ser	Lys	Val	Arg	Ile	Glu	Ala	Cys
			180					185					190		
Met	Trp	Gly	Ile	Gly	Thr	Ala	Ile	Gly	Glu	Leu	Pro	Pro	Tyr	Phe	Met
		195					200					205			
Ala	Arg	Ala	Λla	Arg	Leu	Ser	Gly	Ala	Glu	Pro	Asp	Asp	Glu	Glu	Tyr
	210					215					220				
Gln	Glu	Phe	Glu	Glu	Met	Leu	Glu	His	Ala	Glu	Ser	Ala	Gln	Asp	Phe
225					230					235					240
Ala	Ser	Arg	Ala	Lys	Ļeu	Ala	Val	G1n	Lys	Leu	Val	Gln	Lys	Val	Gly
				245					250					255	
Phe	Phe	Gly	Ile	Leu	Ala	Cys	Ala	Ser	Ile	Pro	Asn	Pro	Leu	Phe	Asp
			260					265					270		
Leu	Ala	Gly	Ile	Thr	Cys	Gly	His	Phe	Leu	Val	Pro	Phe	Trp	Thr	Phe
		275					280					285	,		
Pha	Glv	Δla	Thr	ررم ا	Ile	Glv	l.vs	Ala	Tle	Tle	Lvs	Met	His	Ile	Glr

	290					295					300				
Lys	Ile	Phe	Val	Ile	Ile	Thr	Phe	Ser	Lys	His	Ile	Val	Glu	G1n	Met
305					310					315					320
Val	Ala	Phe	Ile	Gly	Ala	Val	Pro	Gly	Ile	Gly	Pro	Ser	Leu	G1n	Lys
				325					330					335	
Pro	Phe	Gln	Glu	Tyr	Leu	Glu	Ala	Gln	Arg	Gln	Lys	Leu	His	His	Lys
			340					345					350		
Ser	Glu	Met	Gly	Thr	Pro	Gln	Gly	Glu	Asn	Trp	Leu	Ser	Trp	Met	Phe
		355					360					365			
Glu	Lys	Leu	Val	Val	Val	Met	Val	Cys	Tyr	Phe	Ile	Leu	Ser	Ile	He
	370					375					380				
Asn	Ser	Met	Ala	Gln	Ser	Tyr	Ala	Lys	Arg	Ile	Gln	Gḷn	Arg	Leu	Asn
385					390					395					400
Ser	Glu	Glu	Lys	Thr	Lys										
				405											
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<21	1> 4	53													
<21	2> P	ŖT			. ,										
<21	3> H	omo	sapi	ens											
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ı				5					10					15	
Thr	Gln	Ala	Val	Ser	Lys	Leu	Trp	Val	Pro	Asn	Thr	Asp	Phe	Asp	Val
			20					25					30		

Ala	Ala	Asn	Trp	Ser	Gln	Asn	Arg	Thr	Pro	Cys	Ala	Gly	Gly	Ala	Val
		35					40					45			
Glu	Phe	Pro	Ala	Asp	Lys	Met	Val	Ser	Val	Leu	Val	Gln	Glu	Gly	His
	50					55					60				
Ala	Val	Ser	Asp	Met	Leu	Leu	Pro	Leu	Asp	Gly	Glu	Leu	Val	Leu	Ala
65					70					75					80
Ser	Gly	Ala	Gly	Phe	Gly	Val	Ser	Asp	Val	Gly	Ser	His	Leu	Asp	Cys
				85					90					95	
Gly	Ala	Gly	Glu	Pro	Ala	Val	Phe	Arg	Asp	Ser	Asp	Arg	Phe	Ser	Trp
			100					105					110		
His	Asp	Pro	His	Leu	Trp	Arg	Ser	Gly	Asp	Glu	Ala	Pro	Gly	Leu	Phe
		115					120					125			
Phe	Val	Asp	Ala	Glu	Arg	Val	Pro	Cys	Arg	His	Asp	Asp	Val	Phe	Phe
	130					135					140				
Pro	Pro	Ser	Ala	Ser	Phe	Arg	Val	Gly	Leu	Gly	Pro	Gly	Ala	Ser	Pro
145					150					155					160
Val	Arg	Val	Arg	Ser	Ile	Ser	Ala	Leu	Gly	Arg	Thr	Phe	Thr	Arg	Asp
				165					170					175	
Glu	Asp	Leu	Ala	Val	Phe	Leu	Ala	Ser	Arg	Ala	Gly	Arg	Leu	Arg	Phe
			180					185					190		
His	Gly	Pro	Gly	Ala	Leu	Ser	Val	Gly	Pro	Glu	Asp	Cys	Ala	Asp	Pro
		195					200					205			
Ser	Gly	Cys	Val	Cys	Gly	Asn	Ala	Glu	Ala	Gln	Pro	Trp	Ile	Cys	Ala
	210					215					220				
Ala	ادم آ	יים ו	Gln	Pro	Leu	Glv	Glv	Arø	Cvs	Pro	Gln	Ala	Ala	Cys	His

225					230					235					240
Ser	Ala	Ļeu	Arg	Pro	Gln	Gly	Gln	Cys	Cys	Asp	Leu	Cys	Gly	Ala	Val
				245			•		250					255	
Val	Leu	Leu	Thr	His	Gly	Pro	Ala	Phe	Asp	Leu	Glu	Arg	Tyr	Arg	Ala
			260					265					270		
Arg	Ile	Leu	Asp	Thr	Phe	Leu	Gly	Leu	Pro	Gln	Tyr	His	Gly	Leu	Gln
		275					280					285			
Val	Ala	Val	Ser	Lys	Val	Pro	Arg	Ser	Ser	Arg	Leu	Arg	Glu	Ala	Asp
	290					295					300				
Thr	Glu	Ile	Gln	Val	Val	Leu	Val	Glu	Asn	Gly	Pro	Glu	Thr	Gly	Gly
305					310					315					320
Ala	Gly	Arg	Leu	Åla	Arg	Ala	Leu	Leu	Ala	Asp	Val	Ala	Glu	Asn	Gly
				325					330					335	
Glu	Ala	Leu	Gly	Val	Leu	Glu	Ala	Thr	Met	Arg	Glu	Ser	Gly	Ala	His
			340)				345					350		
Val	Trp	Gly	Ser	Ser	Ala	Ala	Gly	Leu	Ala	Gly	Gly	Val	Ala	Ala	Ala
		355	;				360	•				365			
Val	Leu	ı Let	ı Ala	. Leu	Leu	Val	Leu	Leu	Val	Ala	Pro	Pro	Leu	Leu	Arg
	370)				375	5				380)-			
Arg	, Ala	a Gly	/ Arg	g Leu	ı Arg	Trp	Arg	Arg	His	s Glu	ı Ala	Ala	Ala	Pro	Ala
385	5				390)				395	5				400
Gly	/ Ala	a Pro	Lei	ı Gly	, Phe	Arg	g Asr	Pro	Va]	l Phe	e Asp	Val	Thr	Ala	Ser
				408	5				410)				415	5
Glu	ı Glı	u Lei	u Pro	o Lei	u Pro	Arı	g Arg	z Lei	ı Sei	r Lei	ı Val	l Pro	Lys	s Ala	a Ala
			420	0				42	5				430)	

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Ala Asp Ser Thr Ser His Ser Tyr Phe Val Asn Pro Leu Phe Ala Gly

435

440

445

Ala Glu Ala Glu Ala

450

⟨210⟩ 126

<211> 59

<212> PRT

<213> Homo sapiens

<400> 126

Met Thr Ser Val Ser Thr Gln Leu Ser Leu Val Leu Met Ser Leu Leu

1 5 10 15

Leu Val Leu Pro Val Val Glu Ala Val Glu Ala Gly Asp Ala Ile Ala

20 25 30

Leu Leu Leu Gly Val Val Leu Ser Ile Thr Gly Ile Cys Ala Cys Leu

35 40 45

Gly Val Tyr Ala Arg Lys Arg Asn Gly Gln Met

50 55

<210> 127

<211> 210

<212> PRT

<213> Homo sapiens

<400> 127

Met Ala Leu Pro Gln Met Cys Asp Gly Ser His Leu Ala Ser Thr Leu

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Arg	Tyr	Cys	Met	Thr	Val	Ser	Gly	Thr	Val	Val	Leu	Val	Ala	Gly	Thr
			20					25					30		
Leu	Cys	Phe	Ala	Trp	Trp	Ser	Glu	Gly	Asp	Ala	Thr	Ala	G ln	Pro	Gly
		35					40					45			
Gln	Leu	Ala	Pro	Pro	Thr	Glu	Tyr	Pro	Val	Pro	Glu	Gly	Pro	Ser	Pro
	50					55					60				
Leu	Leu	Arg	Ser	Val	Ser	Phe	Val	Cys	Cys	Gly	Ala	Gly	Gly	Leu	Leu
65					70					75					80
Leu	Leu	Ile	Gly	Leu	Leu	Trp	Ser	Val	Lys	Ala	Ser	Ile	Pro	Gly	Pro
				85					90					95	
Pro	Arg	Trp	Asp	Pro	Tyr	His	Leu	Ser	Arg	Asp	Leu	Tyr	Tyr	Leu	Thr
			100				•	105					110		
Val	Glu	Ser	Ser	Glu	Lys	Ģlu	Ser	Cys	Arg	Thr	Pro	Lys	Val	Val	Asp
		115					120					125			
Ile	Pro	Thr	Tyr	Glu	Glu	Ala	Val	Ser	Phe	Pro	Val	Ala	Glu	Gly	Pro
	130					135					140				
Pro	Thr	Pro	Pro	Ala	Tyr	Pro	Thr	Glu	Glu	Ala	Leu	Glu	Pro	Ser	Gly
145					150					155					160
Ser	Arg	Asp	Ala	Leu	Leu	Ser	Thr	Gln	Pro	Ala	Trp	Pro	Pro	Pro	Ser
				165					170					175	
Tyr	Glu	Ser	Ile	Ser	Leu	Ala	Leu	Asp	Ala	Val	Ser	Ala	Glu	Thr	Thr
			180					185					190		
Pro	Ser	Ala	Thr	Arg	Ser	Cys	Ser	Gly	Leu	Val	Gln	Thr	Ala	Arg	Gly
•		105					200					205			

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Gly Ser <210> 128 <211> 165 <212> PRT <213> Homo sapiens <400> 128 Met Asp Ser Ser Arg Ala Arg Gln Gln Leu Arg Arg Phe Leu Leu Leu Pro Asp Ala Glu Ala Gln Leu Asp Arg Glu Gly Asp Ala Gly Pro Glu Thr Ser Thr Ala Val Glu Lys Lys Glu Lys Pro Leu Pro Arg Leu Asn Ile His Ser Gly Phe Trp Ile Leu Ala Ser Ile Val Val Thr Tyr

Tyr Val Asp Phe Phe Lys Thr Leu Lys Glu Asn Phe His Thr Ser Ser

Trp Phe Leu Cys Gly Ser Ala Leu Leu Leu Val Ser Leu Ser Ile Ala

Phe Tyr Cys Ile Val Tyr Leu Glu Trp Tyr Cys Gly Ile Gly Glu Tyr

Asp Val Lys Tyr Pro Ala Leu Ile Pro Ile Thr Thr Ala Ser Phe Ile

Ala Ala Gly Ile Cys Phe Asn Ile Ala Leu Trp His Val Trp Ser Phe

	130					135					140				
Phe '	Thr	Pro	Leu	Leu	Leu	Phe	Thr	Gln	Phe	Met _.	Gly	Val	Val	Met	Phe
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Ile	Thr	Leu	Leu	Gly											
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Met	Leu	Gln	Thr	Ser	Asn	Tyr	- Ser	Leu	ı Val	Leu	Ser	Leu	Gln	Phe	Leu
l				5	;				10)				15	•
Leu	Leu	Ser	Туз	r Asp	Leu	Phe	e Val	Ası	n Sei	. Phe	Ser	Glu	Leu	Leu	Gln
			20					2					30		
Lys	Thr	Pro	Va.	l Ile	Glr	ı Lei	ı Val	Le	u Phe	e Ile	Ile	Gln	Asp	Ile	Ala
		35	5				4()				45	ì		
Val	Leu	Phe	e Asi	n Ile	e Ile	e Ile	e Ile	e Ph	e Le	u Met	Phe	Phe	Asr	Thr	. Phe
	50) .				5	5				60)	•		
Val	Phe	Gli	n Al	a Gl	y Le	u Va	l Ası	n Le	u Le	u Phe	His	Lys	Phe	e Lys	s Gly
65					7	0				75	•				80
Thr	Ile	: I1	e Le	u Th	r Al	a Va	l Ty	r Ph	e Al	a Leu	ı Ser	· Ile	e Se		u His
				8					9					9	
Va]	Trp) Va	l Me	t As	n Le	u Ar	g Tr	рĹу	s As	n Sei	. Asr	ı Sei	r Ph	e Il	e Trp
			10	0				10	5				11	0	

Thr	As	р (Gly	Leu	Gln	Met	Leu	Phe	Val	Phe	Gln	Arg	Leu	Ala	Ala	Val
		1	115					120					125			
Leu	Ty.	r (Cys	Tyr	Phe	Tyr	Lys	Arg	Thr	Ala	Val	Arg	Leu	Gly	Asp	Pro
	13	0					135					140				
His	Ph	e '	Tyr	G1n	Asp	Ser	Leu	Trp	Leu	Arg	Lys	Glu	Phe	Met	Gln	Val
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Arg	Ar	g														
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]	l					5				10)				15	5
Sei	r A	rg	Tyr	- Gl	n Gli	n Lei	ı Glr	n Asn	Glu	ı Glu	ı Glu	ı Sei	r Gly	y Glu	ı Pro	o Glu
				2	0				25	5				30)	
Gli	n A	la	Ala	a G1	y As	p Ala	a Pro	Pro	Pro	о Туі	r Se	r Se	r Il	e Se	r Ala	a Glu
			38	5				40)				4	5		
Se	r A	la	Ala	а Ту	r Ph	e As	р Ту	r Lys	s Ası	p Gl	u Se	r Gl	y Ph	e Pr	o Ly	s Pro
		50					5	5				6	0			
Pr	o S	er	Тy	r As	n Va	1 Al	a Th	r Th	r Le	u Pr	o Se	r Ty	r As	p Gl	u Al	a Glu
6	5					7	0				7	5				80
Ar	g T	hr	Ly	s Al	a Gl	u Al	a Th	r Il	e Pr	o Le	u Va	1 Pr	o Gl	y Ar	g As	p Glu
					8	5				9	0				9	5

Asp	Phe	Val	Gly	Arg	Asp	Asp	Phe	Asp	Asp	Ala	Asp	Gln	Leu	Arg	Ile	
			100					105					110	•	•	
Gly	Asn	Asp	Gly	Ile	Phe	Met	Leu	Thr	Phe	Phe	Met	Ala	Phe	Leu	Phe	
		115					120					125				
Asn	Trp	Ile	Gly	Phe	Phe	Leu	Ser	Phe	Cys	Leu	Thr	Thr	Ser	Ala	Ala	
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Gly	Arg	Tyr	Gly	Ala	Ile	Ser	Gly	Phe	Gly	Leu	Ser	Leu	Ile	Lys	Trp	
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				165					170					175		
Gln	Tyr	Trp	Leu	Trp	Trp	Val	Phe	Leu	Val	Leu	Gly	Phe	Leu	Leu	Phe	
			180					185				•	190			
Leu	Arg	Gly	Phe	Ile	Asn	Tyr	Ala	Lys	Val	Arg	Lys	Met	Pro	Glu	Thr	
		195					200					205	i			
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ato	gtgo	ttg	tcaa	caag	gc g	gctgo	tgad	c a	ccta	eggtt	tco	ccgt	cacc	aati	tttcctt	180

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g	gaattggac	agatggcagc	caccataatg	atactatatg	tgtccaagct	aaacaaaatc	240
a	ttcacttcc	ctgattttga	taagaaaatt	cctgtaaagc	tgtttcctct	gcctctcctc	300
ta	acgttggaa	accacataag	tggattatca	agcacaagta	aattaagcct	accgatgttc	360
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a	aatacggag	tacttttcta	caatgcctgc	ttcatgatta	tcccaactct	tattattagt	660
g	tctccactg	gagacctgca	acaggctact	gaattcaacc	aatggaagaa	tgttgtgttt	720
a	tcctacagt	ttcttctttc	ctgttttttg	gggtttctgc	tgatgtactc	cacggttctg	780
t	gcagctatt	acaattcagc	cctgacgaca	gcagtggttg	gagccatcaa	gaatgtatcc	840
g	ttgcctaca	ttgggatatt	aatcggtgga	gactacattt	tctctttgtt	aaactttgta	900
g	ggttaaata	tttgcatggc	agggggcttg	agatattcct	tttaacact	gagcagccag	960
t	taaaaccta	aacctgtggg	tgaagaaaac	atctgtttgg	atttgaagag	С	1011

<210> 132

<211> 708

<212> DNA

<213> Homo sapiens

<400> 132

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ctcgcattaa	tgctgcaaag	agactctgca	gatagtgaaa	aagatggagg	gccccaattt	360
gcagaggtgt	ttgtcattgt	ctggtttggt	gcagttacca	tcaccctcaa	ctcaaaactt	420
cttggaggga	acatatcttt	ttttcagagc	ctctgtgtgc	tgggttactg	tatacttccc	480
ttgacagtag	caatgctgat	ttgccggctg	gtacttttgg	ctgatccagg	acctgtaaac	540
ttcatggttc	ggctttttgt	ggtgattgtg	atgtttgcct	ggtctatagt	tgcctccaca	600
gctttccttg	ctgatagcca	gcctccaaac	cgcagagccc	tagctgttta	tcctgttttc	660
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<210> 133

<211> 1680

<212> DNA

(213) Homo sapiens

⟨400⟩ 133

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tgcatcatcc	acggggtgct	gcagatcctg	ttccaggccg	tcctcatcgt	cagcgggaac	900
ctcagcttcc	tgaactggct	gactatggtg	cccagcctgg	cctgctttga	tgacgccacc	960
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<210> 134

<211> 1218

<212> DNA

<213> Homo sapiens

<400> 134

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aggcagaata ttgtcctgtg gagacagccg ctcattacct tgcagtattt ttctctggaa 180

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atccttgtaa	tcttgaagga	atggacctca	aaattatggc	atcgtcaaag	cattgtggtg	240
tctttttac	tgctgcttgc	tgtgcttata	gctacgtatt	atgttgaagg	agtgcatcaa	300
cagtatgtgc	aacgtataga	gaaacagttt	cttttgtatg	cctactggat	aggcttagga	360
attttgtctt	ctgttgggct	tggaacaggg	ctgcacacct	ttctgcttta	tctgggtcca	420
catatagcct	cagttacatt	agctgcttat	gaatgcaatt	cagttaattt	tcccgaacca	480
ccctatcctg	atcagattat	ttgtccagat	gaagagggca	ctgaaggaac	catttctttg	540
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gatgaagagt	atcaggaatt	tgaagagatg	ctggaacatg	cagagtctgc	acaagacttt	720
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atgcatatco	agaaaattt	tgttataata	acattcagca	agcacatagt	ggagcaaatg	960
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<210> 135

<211> 1359

<212> DNA

<213> Homo sapiens

⟨400⟩ 135

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caagaaggtc	acgccgtctc	agacatgctc	ctgccgctgg	atggggaact	cgtcctggct	240
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⟨210⟩ 136

<211> 177

275/307

<212> DNA

<213> Homo sapiens

<400> 136

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atg	acctcag	tttcaacaca	gttgtcctta	gtcctcatgt	cactgctttt	ggtgctgcct	60
gtt	gtggaag	cagtagaagc	cggtgatgca	atcgcccttt	tgttaggtgt	ggttctcagc	120
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<210> 137

<211> 630

<212> DNA

<213> Homo sapiens

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gtattccaga	gactagcagc	agtgttgtac	tgctacttct	ataaacggac	agccgtaaga	420
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⟨210⟩ 140

<211> 663

<212> DNA

<213> Homo sapiens

<400> 140

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ccttacagca	gcatttctgc	agagagcgca	gcatattttg	actacaagga	tgagtctggg	180
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210> 141	
211> 1622	
212> DNA	
213> Homo sapiens	
220>	
221> CDS	
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(400> 141	
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Met Thr Ala Gly Gly Gln Ala Glu Ala Glu Gly	
1 5 10	
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Ala Gly Gly Glu Pro Gly Ala Ala Arg Leu Pro Ser Arg Val Ala Arg	
15 20 25	
ctg ctg tcg gcg ctc ttc tac ggg acc tgc tcc ttc ctc atc gtg ctt	206
Leu Leu Ser Ala Leu Phe Tyr Gly Thr Cys Ser Phe Leu Ile Val Leu	
30 35 40	
gtc aac aag gcg ctg ctg acc acc tac ggt ttc ccg tca cca att ttc	254
Val Asn Lys Ala Leu Leu Thr Thr Tyr Gly Phe Pro Ser Pro Ile Phe	
45 50 55	
ctt gga att gga cag atg gca gcc acc ata atg ata cta tat gtg tcc	302
Leu Gly Ile Gly Gln Met Ala Ala Thr Ile Met Ile Leu Tyr Val Ser	
60 65 70 75	
and otherwise are att cac tto cot gat ttt gat aag aaa att cot	350

Lys	Leu	Asn	Lys	Ile	Ile	His	Phe	Pro	Asp	Phe	Asp	Lys	Lys	Ile	Pro	
				80					85					90		
gta	aag	ctg	ttt	cct	ctg	cct	ctc	ctc	tac	gtt	gga	aac	cac	ata	agt	398
Val	Lys	Leu	Phe	Pro	Leu	Pro	Leu	Leu	Tyr	Val	Gly	Asn	His	Ile	Ser	
			95					100					105			
gga	tta	tca	agc	aca	agt	aaa	tta	agc	cta	ccg	atg	ttc	acc	gtg	ctc	446
Gly	Leu	Ser	Ser	Thr	Ser	Lys	Leu	Ser	Leu	Pro	Met	Phe	Thr	Val	Leu	
		110					115					120				
agg	aaa	ttc	acc	att	cca	ctt	acc	tta	ctt	ctg	gaa	acc	atc	ata	ctt	494
Arg	Lys	Phe	Thr	Ile	Pro	Leu	Thr	Leu	Leu	Leu	Glu	Thr	Ile	Ile	Leu	
	125					130					135					
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Gly	Lys	Gln	Tyr	Ser	Leu	Asn	Ile	Ile	Leu	Ser	Val	Phe	Ala	Ile	Ile	
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ctc	ggg	gct	ttc	ata	gca	gct	ggg	tct	gac	ctt	gct	ttt	aac	tta	gaa	590
Leu	Gly	Ala	Phe	Ile	Ala	Ala	Gly	Ser	Asp	Leu	Ala	Phė	Asn	Leu	Glu	
				160					165					170)	
ggc	tat	att	ttt	gta	ttc	ctg	aat	gat	atc	ttc	aca	gca	gca	aat	gga	638
Gly	Tyr	·Ile	Phe	Val	Phe	Leu	Asn	Asp	Ile	Phe	Thr	Ala	Ala	Asn	Gly	
			175	ı				180					185	I		
gtt	tat	acc	aaa	cag	aaa	atg	gac	cca	aag	gag	cta	ggg	aaa	tac	gga	686
Va J	Tyr	Thr	Lys	Gln	Lys	Met	Asp	Pro	Lys	Glu	l Leu	Gly	Lys	Tyr	Gly	
		190)				195	,				200)			
gta	cti	ttc	tac	aat	gcc	tgc	ttc	atg	att	ato	cca	act	ctt	att	att	734
Val	Leu	ı Phe	Tvr	Asn	Ala	Cys	Phe	e Met	Ile	· Ile	e Pro	Thr	· Leı	ı Ile	lle	

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	205					210					215					
agt	gtc	tcc	act	gga	gac	ctg	caa	cag	gct	act	gaa	tţc	aac	caa	tgg	782
Ser	Val	Ser	Thr	Gly	Asp	Leu	Gln	Gln	Ala	Thr	Glu	Phe	Asn	Gln	Trp	•
220					225					230					235	
aag	aat	gtt	gtg	ttt	atc	cta	cag	ttt	ctt	ctt	tcc	tgt	ttt	ttg	ggg	830
Lys	Asn	Val	Val	Phe	Ile	Leu	Gln	Phe	Leu	Leu	Ser	Cys	Phe	Leu	Gly	
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Phe	Leu	Leu	Met	Tyr	Ser	Thr	Val	Leu	Cys	Ser	Tyr	Tyr	Asn	Ser	Ala	
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ctg	acg	aca	gca	gtg	gtt	gga	gcc	atc	aag	aat	gta	tcc	gtt	gcc	tac	926
Leu	Thr	Thr	Ala	Val	Val	Gly	Ala	Ile	Lys	Asn	Val	Ser	Val	Λla	Tyr	
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att	ggg	ata	tta	atc	ggt	gga	gac	tac	att	ttc	tct	ttg	tta	aac	ttt	974
Ile	Gly	Ile	Leu	Ile	Gly	Gly	Asp	Tyr	Ile	Phe	Ser	Leu	Leu	Asn	Phe	
	285					290					295					
gta	ggg	tta	aat	att	tgc	atg	gca	ggg	ggc	ttg	aga	tat	tcc	ttt	tta	1022
Val	Gly	Leu	Asn	Ile	Cys	Met	Ala	Gly	Gly	Leu	Arg	Tyr	Ser	Phe	Leu	
300					305					310				٠	315	
aca	ctg	agc	agc	cag	tta	aaa	cct	aaa	cct	gtg	ggt	gaa	gaa	aac	atc	1070
Thr	Leu	Ser	Ser	Gln	Leu	Lys	Pro	Lys	Pro	Val	Gly	Glu	Glu	Asn	Ile	
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tgt	ttg	gat	ttg	aag	agc	ta	aaga	gtct	gc a	gcag	gatt	g ga	gact	gact		1120
Cys	Leu	Asp	Leu	Lys	Ser											

335

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Ser Pro Gly Asp Pro Gly Thr Ala Ser Pro Arg Pro Leu Phe Ala Gly	
10 15 20	

ctt tca gat ata tcc atc tca caa gac atc ccc gta gaa gga gaa atc 149

Leu	Ser	Asp	Ile	Ser	Ile	Ser	Gln	Asp	Ile	Pro	Val	Glu	Gly	Glu	Ile	
		25					30					35				
acc	att	cct	atg	aga	tct	cgc	atc	cgg	gag	ttt	gac	agc	tcc	aca	tta	197
Thr	Ile	Pro	Met	Arg	Ser	Arg	Ile	Arg	Glu	Phe	Asp	Ser	Ser	Thr	Leu	
	40					45					50					
aat	gaa	tct	gtt	cgc	aat	acc	atc	atg	cgt	gat	cta	aaa	gct	gtt	ggg	245
Asn	Glu	Ser	Val	Arg	Asn	Thr	Ile	Met	Arg	Asp	Leu	Lys	Ala	Val	Gly	
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aaa	aaa	ttc	atg	cat	gtt	ttg	tac	cca	agg	aaa	agt	aat	act	ctt	ttg	293
Lys	Lys	Phe	Met	His	Val	Leu	Tyr	Pro	Arg	Lys	Ser	Asn	Thr	Leu	Leu	
				75					80					85		
aga	gat	tgg	gat	ttg	tgg	ggc	cct	ttg	atc	ctt	tgt	gtg	aca	ctc	gca	341
Arg	Asp	Trp	Asp	Leu	Trp	Gly	Pro	Leu	Ile	Leu	Cy.s	Val	Thr	Leu	Ala	
			90					95					100			
tta	atg	ctg	caa	aga	gac	tct	gca	gat	agt	gaa	aaa	gat	gga	ggg	ccc	389
Leu	Met	Leu	Gln	Arg	Asp	Ser	Ala	Asp	Ser	Glu	Lys	Asp	Gly	Gly	Pro	
		105					110					115				
caa	ttt	gca	gag	gtg	ttt	gtc	att	gtc	tgg	ttt	ggt	gca	gtt	acc	atc	437
Gln	Phe	Ala	Glu	Val	Phe	Val	Ile	Val	Trp	Phe	Gly	Ala	Val	Thr	Ile	
	120					125					130					
acc	ctc	aac	tca	aaa	ctt	ctt	gga	ggg	aac	ata	tct	ttt	ttt	cag	agc	485
Thr	Leu	Asn	Ser	Lys	Leu	Leu	Gly	Gly	Asn	Ile	Ser	Phe	Phe	Gln	Ser	
135					140					145					150	
ctc	tgt	gtg	ctg	ggt	tac	tgt	ata	ctt	ccc	ttg	aca	gta	gca	atg	ctg	533
1 011	Cvc	Val	Lau	Clv	Tyr	Cve	T۱۵	يرم ا	Pro	ينم ا	Thr	Val	Ala	Met	Leu	

155 160 165	
att tgc cgg ctg gta ctt ttg gct gat cca gga cct gta aac ttc atg	581
Ile Cys Arg Leu Val Leu Leu Ala Asp Pro Gly Pro Val Asn Phe Met	
170 175 180	
gtt cgg ctt ttt gtg gtg att gtg atg ttt gcc tgg tct ata gtt gcc	629
Val Arg Leu Phe Val Val Ile Val Met Phe Ala Trp Ser Ile Val Ala	
185 190 195	
tcc aca gct ttc ctt gct gat agc cag cct cca aac cgc aga gcc cta	677
Ser Thr Ala Phe Leu Ala Asp Ser Gln Pro Pro Asn Arg Arg Ala Leu	
200 205 210	
gct gtt tat cct gtt ttc ctg ttt tac ttt gtc atc agt tgg atg att	725
Ala Val Tyr Pro Val Phe Leu Phe Tyr Phe Val Ile Ser Trp Met Ile	
215 220 225 230	
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Leu Thr Phe Thr Pro Gln	
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cgg aag act g	gg tac tcg ga	at ccg gag cct	gag tcg ccg ccc	gcg ccg 98
Arg Lys Thr G	ly Tyr Ser As	sp Pro Glu Pro	Glu Ser Pro Pro	Ala Pro
	15	20		25
ggg cgt ggc c	cc gca ggc to	ct ccg gcc cat	ctc cac acg ggc	acc ttc 146
Gly Arg Gly P	ro Ala Gly Se	er Pro Ala His	Leu His Thr Gly	Thr Phe
:	30	35	40	
tgg ctg acc c	gg atc gtg ct	c ctg aag gcc	cta gcc ttc gtg	tac ttc 194
Trp Leu Thr A	rg Ile Val Le	eu Leu Lys Ala	Leu Ala Phe Val	Tyr Phe
45		50	55	
gtg gca ttc c	tg gtg gct tt	c cat cag aac	aag cag ctc atc	ggt gac 242
Val Ala Phe L	eu Val Ala Ph	e His Gln Asn	Lys Gln Leu Ile	Gly Asp
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agg ggg ctg c	tt ccc tgc ag	a gtg ttc ctg	aag aac ttc cag	cag tac 290
Arg Gly Leu Lo	eu Pro Cys Ar	g Val Phe Leu	Lys Asn Phe Gln	Gln Tyr
75	80		85	90
ttc cag gac ag	gg acg agc tg	g gaa gtc ttc	agc tac atg ccc	acc atc 338

95 100 105

Phe Gln Asp Arg Thr Ser Trp Glu Val Phe Ser Tyr Met Pro Thr Ile

ctc	tgg	ctg	atg	gac	tgg	tca	gac	atg	aac	tcc	aac	ctg	gac	ttg	ctg	386
Leu	Trp	Leu	Met	Asp	Trp	Ser	Asp	Met	Asn	Ser	Asn	Leu	Asp	Leu	Leu	
			110					115					120			
gct	ctt	ctc	gga	ctg	ggc	atc	tcg	tct	ttc	gta	ctg	atc	acg	ggc	tgc	434
Ala	Leu	Leu	Gly	Leu	Gly	Ile	Ser	Ser	Phe	Val	Leu	Ile	Thr	Gly	Cys	
		125					130					135				
gcc	aac	atg	ctt	çtc	atg	gct	gcc	ctg	tgg	ggc	ctc	tac	atg	tcc	ctg	482
Ala	Asn	Met	Leu	Leu	Met	Ala	Ala	Leu	Trp	Gly	Leu	Tyr	Met	Ser	Leu	
	140					145					150					
gtt	aat	gtg	ggc	cat	gtc	tgg	tac	tct	ttc	gga	tgg	gag	tcc	cag	ctt	530
Val	Asn	Val	Gly	His	Val	Trp	Tyr	Ser	Phe	Gly	Trp	Glu	Ser	Gln	Leu	
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ctg	gag	acg	ggg	ttc	ctg	ggg	atc	ttc	ctg	tgc	cct	ctg	tgg	acg	ctg	578
Leu	Glu	Thr	Gly	Phe	Leu	Gly	Ile	Phe	Leu	Cys	Pro	Leu	Trp	Thr	Leu	
				175					180				•	185		
tca	agg	ctg	ccc	cag	cat	acc	ccc	aca	tcc	cgg	att	gtc	ctg	tgg	ggc	626
Ser	Arg	Leu	Pro	G1n	His	Thr	Pro	Thr	Ser	Arg	Ile	Val	Leu	Trp	Gly	
			190					195					200			
ttc	cgg	tgg	ctg	atc	ttc	agg	atc	atg	ctt	gga	gca	ggc	ctg	atc	aag	674
Phe	Arg	Trp	Leu	Ile	Phe	Arg	Ile	Met	Leu	Gly	Ala	Gly	Leu	Ile	Lys	
		205					210					215				
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Ile	Arg	Gly	Asp	Arg	Cys	Trp	Arg	Asp	Leu	Thr	Cys	Met	Asp	Phe	His	
	220					225					230					
tat	asa	200		cca	ato	ccc	aat	cet	øte	gca	tac	tac	ctg	cac	cac	770

Tyr	Glu	Thr	Gln	Pro	Met	Pro	Asn	Pro	Val	Ala	Tyr	Tyr	Leu	His	His	
235					240					245					250	
tca	ccc	tgg	tgg	ttc	cat	cgc	ttc	gag	acg	ctc	agc	aac	cac	ttc	atc	818
Ser	Pro	Trp	Trp	Phe	His	Arg	Phe	Glu	Thr	Leu	Ser	Asn	His	Phe	Ile	
				255					260					265		
gag	ctc	ctg	gtg	ccc	ttc	ttc	ctc	ttc	ctc	ggc	cgg	cgg	gcg	tgc	atc	866
Glu	Leu	Leu	Val	Pro	Phe	Phe	Leu	Phe	Leu	Gly	Arg	Arg	Ala	Cys	Ile	
			270					275					280			
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Ile	His	Gly	Val	Leu	Gln	Ile	Leu	Phe	Gln	Ala	Val	Leu	Ile	Val	Ser	
		285					290					295				
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	300					305					310					
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Cys	Phe	Asp	Asp	Ala	Thr	Leu	Gly	Phe	Leu	Phe	Pro	Ser	Gly	Pro	Gly	
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Ser	Leu	Lys	Asp	Arg	Val	Leu	Gln	Met	Gln	Arg	Asp	Iļe	Arg	Gly	Ala	
				335					340					345		
cgg	ccc	gag	ccc	aga	ttc	ggc	tcc	gtg	gtg	cgg	cgt	gca	gcc	aac	gtc	1106
Arg	Pro	Glu	Pro	Arg	Phe	Gly	Ser	Val	Val	Arg	Arg	Ala	Ala	Asn	Val	
			350					355					360			
tcg	ctg	ggc	gtc	ctg	ctg	gcc	tgg	ctc	agc	gtg	ссс	gtg	gtc	ctc	aac	1154
Ser	Leu	Gly	Val	Leu	Leu	Ala	Trp	Leu	Ser	Val	Pro	Val	Val	Leu	Asn	

		365					370					375				
ttg	ctg	agc	tcc	agg	cag	gtc	atg	aac	acc	cac	ttc _.	aac	tct	ctt _.	cac	1202
Leu	Leu	Ser	Ser	Arg	Gln	Val	Met	Asn	Thr	His	Phe	Asn	Ser	Leu	His	
	380					385					390					
atc	gtc	aac	act	tac	ggg	gcc	ttc	gga	agc	atc	acc	aag	gag	cgg	gcg	1250
Ile	Val	Asn	Thr	Tyr	Gly	Ala	Phe	Gly	Ser	Ile	Thr	Lys	Glu	Arg	Ala	
395					400					405					410	
gàg	gtg	atc	ctg	cag	ggc	aca	gcc	agc	tcc	aac	gcc	agc	gcc	ccc	gat	1298
Glu	Val	Ile	Leu	Gln	Gly	Thr	Ala	Ser	Ser	Asn	Ala	Ser	Ala	Pro	Asp	
				415					420					425		
gcc	atg	tgg	gag	gac	tac	gag	ttc	aag	tgc	aag	cca	ggt	gac	ccc	agc	1346
Ala	Met	Trp	Glu	Asp	Tyr	Glu	Phe	Lys	Cys	Lys	Pro	Gly	Asp	Pro	Ser	
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aga	cgg	ccc	tgc	ctc	atc	tcc	ccg	tac	cac	tac	cgc	ctg	gac	tgg	ctg	1394
Arg	Arg	Pro	Cys	Leu	Ile	Ser	Pro	Tyr	His	Tyr	Arg	Leu	Asp	Trp	Leu	
		445					450					455				
atg	tgg	ttc	gcg	gcc	ttc	cag	acc	tac	gag	cac	aac	gac	tgg	atc	atc	1442
Met	Trp	Phe	Ala	Ala	Phe	Gln	Thr	Tyr	Glu	His	Asn	Asp	Trp	Ile	Ile	
	460					465					470					٠
cac	ctg	gct	ggc	aag	ctc	ctg	gcc	agc	gac	gcc	gag	gcc	ttg	tcc	ctg	1490
His	Leu	Ala	Gly	Lys	Leu	Leu	Ala	Ser	Asp	Ala	Glu	Ala	Leu	Ser	Leu	
475					480					485					490	
ctg	gca	cac	aac	ccc	ttc	gcg	ggc	agg	ccc	ccg	ccc	agg	tgg	gtc	cga	1538
Leu	Ala	His	Asn	Pro	Phe	Ala	Gly	Arg	Pro	Pro	Pro	Arg	Trp	Val	Arg	
				495					500	ı				505		

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Gly Glu His Tyr Arg Tyr Lys Phe Ser Arg Pro Gly Gly Arg His Ala	
510 515 520	
gcc gag ggc aag tgg tgg gtg cgg aag agg atc gga gcc tac ttc cct	1634
Ala Glu Gly Lys Trp Trp Val Arg Lys Arg Ile Gly Ala Tyr Phe Pro	
525 530 535	
ccg ctc agc ctg gag gag ctg agg ccc tac ttc agg gac cgt ggg tgg	1682
Pro Leu Ser Leu Glu Glu Leu Arg Pro Tyr Phe Arg Asp Arg Gly Trp	
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Met Ala Glu	

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aat	gga	aaa	aat	tgt	gac	cag	aga	cgt	gta	gca	atg	aac	aag	gaa	cat		163
Asn	Gly	Lys	Asn	Cys	Asṗ	Gln	Arg	Arg	Val	Ala	Met	Asn	Lys	Glu	His		
	5					10					15						
cat	aat	gga	aat	ttc	aca	gac	ccc	tct	tca	gtg	aat	gaa	aag	aag	agg		211
His	Asn	Gly	Asn	Phe	Thr	Asp	Pro	Ser	Ser	Val	Asn	Glu	Lys	Lys	Arg		
20					25					30					35		
agg	gag	cgg	gaa	gaa	agg	cag	aat	att	gtc	ctg	tgg	aga	cag	ccg	ctc		259
Arg	Glu	Arg	Glu	Glu	Arg	Gln	Asn	Ile	Val	Leu	Trp	Arg	Gln	Pro	Leu		
				40					45					50			
att	acc	ttg	cag	tat	ttt	tct	ctg	gaa	atc	ctt	gta	atc	ttg	aag	gaa		307
Ile	Thr	Leu	Gln	Tyr	Phe	Ser	Leu	Glu	Ile	Leu	Val	Ile	Leu	Lys	Glu	•	
			55					60					65				
tgg	acc	tca	aaa	tta	tgg	cat	cgt	caa	agc	att	gtg	gtg	tct	ttt	tta		355
Trp	Thr	Ser	Lys	Leu	Trp	His	Arg	Gln	Ser	Ile	Val	Val	Ser	Phe	Leu		
		70					75					80					
ctg	ctg	ctt	gct	gtg	ctt	ata	gct	acg	tat	tat	gtt	gaa	gga	gtg	cat		403
Ļeu	Leu	Leu	Ala	Val	Leu	Ile	Ala	Thr	Tyr	Tyr	Val	Glu	Gly	Val	His		
	85					90					95						
caa	cag	tat	gtg	caa	cgt	ata	gag	aaa	cag	ttt	ctt	ttg	tat	gcc	tac		451
Gln	Gln	Tyr	Val	Gln	Arg	Ile	Glu	Lys	Gln		Leu	Leu	Tyr	Ala			
100					105					110					115		
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Trp	Ile	Gly	Leu	Gly	Ile	Leu	Ser	Ser		Gly	Leu	Gly	Thr		Leu		
				120					125					130			

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His	Thr	Phe	Leu	Leu	Tyr	Leu	Gly	Pro	His	Ile	Ala	Ser	Val	Thr	Leu	
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Ala	Ala	Tyr	Glu	Cys	Asn	Ser	Val	Asn	Phe	Pro	Glu	Pro	Pro	Tyr	Pro	
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gat	cag	att	att	tgt	cca	gat	gaa	gag	ggc	act	gaa	gga	acc	att	tct	643
Asp	G1n	Ile	Ile	Cys	Pro	Asp	Glu	Glu	Gly	Thr	Glu	Gly	Thr	Ile	Ser	
	165					170					175					
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Leu	Trp	Ser	Ile	Ile	Ser	Lys	Val	Arg	Ile	Glu	Ala	Cys	Met	Trp	Gly	
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atc	ggt	aca	gca	atc	gga	gag	ctg	cct	cca	tat	ttc	atg	gcc	aga	gca	739
Ile	Gly	Thr	Ala	Ile	Gly	Glu	Leu	Pro	Pro	Tyr	Phe	Met	Ala	Arg	Ala	
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gct	cgc	ctc	tca	ggt	gct	gaa	cca	gat	gat	gaa	gag	tat	cag	gaa	ttt	787
Ala	Arg	Leu	Ser	Gly	Ala	Glu	Pro	Asp	Asp	Glu	Glu	Tyr	Gln	Glu	Phe	
			215					220					225			
gaa	gag	atg	ctg	gaa	cat	gca	gag	tct	gca	caa	gac	ttt	gcc	tcc	cgg	835
Glu	Glu	Met	Leu	Glu	His	Ala	Glu	Ser	Ala	Gln	Asp	Phe	Ala	Ser	Arg	
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gcc	aaa	ctg	gca	gtt	caa	aaa	cta	gta	cag	aaa	gtt	gga	ttt	ttt	gga	883
Ala	Lys	Leu	Ala	Val	Gln	Lys	Leu	Val	Gln	Lys	Val	Gly	Phe	Phe	Gly	
	245					250					255					
att	ttg	gcc	tgt	gct	tca	att	cca	aat	cct	tta	ttt	gat	ctg	gct	gga	931

Ile	Leu	Ala	Cys	Ala	Ser	Ile	Pro	Asn	Pro	Leu	Phe	Asp	Leu	Ala	Gly	
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Thr	Leu	Ile	Gly	Lys	Ala	Ile	Ile	Lys	Met	His	Ile	Gln	Lys	Ile	Phe	
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Val	Ile	Ile	Thr	Phe	Ser	Lys	His	Ile	Val	Glu	Gln	Met	Val	Ala	Phe	
		310					315					320				
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Ile	Gly	Ala	Val	Pro	Gly	Ile	Gly	Pro	Ser	Leu	Gln	Lys	Pro	Phe	Gln	
	325					330					335					
gag	tac	ctg	gag	gct	caa	cgg	cag	aag	ctt	cac	cac	aaa	agc	gaa	atg	1171
Glu	Tyr	Leu	Glu	Ala	Gln	Arg	Gln	Lys	Leu	His	His	Lys	Ser	Glu	Met	
340					345					350					355	
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Gly	Thr	Pro	Gln	Gly	Glu	Asn	Trp	Leu	Ser	Trp	Met	Phe	Glu	Lys	Leu	
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gtc	gtt	gtc	atg	gtg	tgt	tac	ttc	atc	cta	tct	atc	att	aac	tcc	atg	1267
Val	Val	Val	Met	Val	Cys	Tyr	Phe	Ile	Leu	Ser	Ile	Ile	Asn	Ser	Met	
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gca	caa	agt	tat	gcc	aaa	cga	atc	cag	cag	cgg	ttg	aac	tca	gag	gag	1315
Ala	Gln	Ser	Tyr	Ala	Lys	Arg	Ile	Gln	Gln	Arg	Leu	Asn	Ser	Glu	Glu	

293/307

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2005

54

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(400) 145

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294/307

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Cys	Arg	His	Asp	Asp	Val	Phe	Phe	Pro	Pro	Ser	Ala	Ser	Phe	Arg	Val	
			140					145					150			
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Gly	Leu	G1y	Pro	Gly	Ala	Ser	Pro	Val	Arg	Val	Arg	Ser	Ile	Ser	Ala	
		155					160					165	•			
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	170					175					180					
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Ser	Arg	Ala	Gly	Arg	Leu	Arg	Phe	His	Gly	Pro	Gly	Ala	Leu	Ser	Val	
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Gly	Pro	Glu	Asp	Cys	Ala	Asp	Pro	Ser	Gly	Cys	Val	Cys	Gly	Asn	Ala	
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Glu	Ala	Gln	Pro	Trp	Ile	Cys	Ala	Ala	Leu	Leu	Gln	Pro	Leu	Gly	Gly	
			220					225					230			
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Arg	Cys	Pro	Gln	Ala	Ala	Cys	His	Ser	Ala	Leu	Arg	Pro	Gln	Gly	Gln	
		235					240					245				
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Cys	Cys	Asp	Leu	Cys	Gly	Ala	Val	Val	Leu	Leu	Thr	His	Gly	Pro	Ala	
	250					255					260					

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Phe	Asp	Leu	Glu	Arg	Tyr	Arg	Ala	Arg	Ile	Leu	Asp	Thr	Phe	Leu	Gly	
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ctg	cct	cag	tac	cac	ggg	ctg	cag	gtg	gcc	gtg	tcc	aag	gtg	cca	cgc	918
Leu	Pro	Gln	Tyr	His	Gly	Leu	G1n	Val	Ala	Val	Ser	Lys	Val	Pro	Arg	
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tcg	tcc	cgg	ctc	cgt	gag	gcc	gat	acg	gag	atc	cag	gtg	gtg	ctg	gtg	966
Ser	Ser	Arg	Leu	Arg	Glu	Ala	Asp	Thr	Glu	Ile	Gln	Val	Val	Leu	Val	
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gag	aat	ggg	ccc	gag	aca	ggc	gga	gcg	ggg	cgg	ctg	gcc	cgg	gcc	ctc	1014
Glu	Asn	Gly	Pro	Glu	Thr	Gly	Gly	Ala	Gly	Arg	Leu	Ala	Arg	Ala	Leu	
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ctg	gcg	gac	gtc	gcc	gag	aac	ggc	gag	gcc	ctc	ggc	gtc	ctg	gag	gcg	1062
Leu	Ala	Asp	Val	Ala	Glu	Asn	Gly	Glu	Ala	Leu	Gly	Val	Leu	Glu	Ala	
	330					335					340					
acc	atg	cgg	gag	tcg	ggc	gca	cac	gtc	tgg	ggc	agc	tcc	gcg	gct	ggg	1110
Thr	Met	Arg	Glu	Ser	Gly	Ala	His	Val	Trp	Gly	Ser	Ser	Ala	Ala	Gly	
345					350					355					360	
ctg	gcg	ggc	ggc	gtg	gcg	gct	gcc	gtg	ctg	ctg	gcg	ctg	ctg	gtc	ctg	1158
Leu	Ala	Gly	Gly	Val	Ala	Ala	Ala	Val	Leu	Leu	Ala	Leu	Leu	Val	Leu	
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ctg	gtg	gcg	ccg	ccg	ctg	ctg	cgc	cgc	gcg	ggg	agg	ctc	agg	tgg	agg	1206
Leu	Val	Ala	Pro	Pro	Leu	Leu	Arg	Arg	Ala	Gly .	Arg	Leu	Arg	Trp	Arg	
			380					385					390			
agg	cac	gag	gcg	gcg	gcc	ccg	gct	gga	gcg	ссс	ctc	ggc	ttc	cgc	aac	1254

Arg His Glu Ala Ala Ala Pro Ala Gly Ala Pro Leu Gly Phe Arg Asn	
395 400 405	
ccg gtg ttc gac gtg acg gcc tcc gag gag ctg ccc ctg ccg cgg cgg	1302
Pro Val Phe Asp Val Thr Ala Ser Glu Glu Leu Pro Leu Pro Arg Arg	
410 415 420	
ctc agc ctg gtt ccg aag gcg gcc gca gac agc acc agc cac agt tac	1350
Leu Ser Leu Val Pro Lys Ala Ala Ala Asp Ser Thr Ser His Ser Tyr	
425 430 435 440	
ttc gtc aac cct ctg ttc gcc ggg gcc gag gcc gag gcc t gagcggccgc	1400
Phe Val Asn Pro Leu Phe Ala Gly Ala Glu Ala Glu Ala	
445 450	
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acctectegt ceageceeca aaccteceet teettteece eteeteegg ggeeaaggae	1520
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ggaaccttca gccctcaaga ttccaacatc atg acc tca gtt	tca aca cag ttg 174
Met Thr Ser Val	Ser Thr Gln Leu
1	5
tcc tta gtc ctc atg tca ctg ctt ttg gtg ctg cct g	tt gtg gaa gca 222
Ser Leu Val Leu Met Ser Leu Leu Val Leu Pro Va	al Val Glu Ala
10 15 20	
gta gaa gcc ggt gat gca atc gcc ctt ttg tta ggt g	tg gtt ctc agc 270
Val Glu Ala Gly Asp Ala Ile Ala Leu Leu Gly Va	al Val Leu Ser
25 30 35	40
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Ile Thr Gly Ile Cys Ala Cys Leu Gly Val Tyr Ala A	rg Lys Arg Asn
45 50	55
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Gly Gln Met	
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gaaatcacca ctgttcggtt ataatcactg cctcctgaat cgttga	aggag tettttaaat 730
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								Me	t Ala	a Le	u Pro	Gl	n Me	t Cy	s Asp	
									l			{	5			
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Gly	Ser	His	Leu	Ala	Ser	Thr	Leu	Arg	Tyr	Cys	Met	Thr	Val	Ser	Gly	
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aca	gtg	gtt	ctg	gtg	gcc	ggg	acg	ctc	tgc	ttc	gct	tgg	tgg	agc	gaa	270
Thr	Val	Val	Leu	Val	Ala	Gly	Thr	Leu	Cys	Phe	Ala	Trp	Trp	Ser	Glu	
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ggg	gat	gca	acc	gcc	cag	cct	ggc	cag	ctg	gcc	cca	ccc	acg	gag	tat	318
Gly	Asp	Ala	Thr	Ala	Gln	Pro	Gly	Gln	Leu	Ala	Pro	Pro	Thr	Glu	Tyr	
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ccg	gtg	cct	gag	ggc	ccc	agc	ccc	ctg	ctc	agg	tcc	gtc	agc	ttc	gtc	366
Pro	Va 1	Pro	Glu	Glv	Pro	Sor	Pro	Leu	Len	Ara	Ser	Val	Ser	Phe	Val	

			60					65					70				
tgc	tgc	ggt	gca	ggt	ggc	ctg	ctg	ctg	ctc	att	ggc	ctg	ctg	tgg	tcc	4	414
Cys	Cys	Gly	Ala	Gly	Gly	Leu	Leu	Leu	Leu	Ile	Gly	Leu	Leu	Trp	Ser		
		75					80					85					
gtc	aag	gcc	agc	atc	cca	ggg	cca	cct	cga	tgg	gac	ccc	tat	cac	ctc	4	162
Val	Lys	Ala	Ser	Ile	Pro	Gly	Pro	Pro	Arg	Trp	Asp	Pro	Tyr	His	Leu		
	90					95					100						
tcc	aga	gac	ctg	tac	tac	ctc	act	gtg	gag	tcc	tca	gag	aag	gag	agc	{	510
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Cys	Arg	Thr	Pro	Lys	Val	Val	Asp	Ile	Pro	Thr	Tyr	Glu	Glu	Ala	Val		
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Ser	Phe	Pro	Val	Ala	Glu	Gly	Pro	Pro	Thr	Pro	Pro	Λla	Tyr	Pro	Thr		
			140					145					150				
gag	gaa	gcc	ctg	gag	cca	agt	gga	tcg	agg	gat	gcc	ctg	ctc	agc	acc	6	554
Glu	Glu	Ala	Leu	Glu	Pro	Ser	Gly	Ser	Arg	Asp	Ala	Leu	Leu	Ser	Thr		
		155					160					165					
cag	ccc	gcc	tgg	cct	cca	ccc	agc	tat	gag	agc	atc	agc	ctt	gct	ctt	7	702
Gln	Pro	Ala	Trp	Pro	Pro	Pro	Ser	Tyr	Glu	Ser	Ile	Ser	Leu	Ala	Leu		
	170					175					180						
gat	gcc	gtt	tct	gca	gag	acg	aca	ccg	agt	gcc	aca	cgc	tcc	tgc	tca	7	'50
Asp	Ala	Val	Ser	Ala	Glu	Thr	Thr	Pro	Ser	Ala	Thr	Arg	Ser	Cys	Ser		
185					190					195					200		

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Gly Leu Val Gln Thr Ala Arg Gly Gly Ser	
205 210	
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gtaggcactc agcaaacgtt cgttgttgaa ggctgttcta tttatctatt gctgtataac	920
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Arg Arg Phe Leu Leu Pro Asp Ala Glu Ala Gln Leu Asp Arg Glu	
15 20 25	
ggt gac gcc ggg ccg gaa acc tcc aca gct gtt gag aaa aag gag aaa	148
Gly Asp Ala Gly Pro Glu Thr Ser Thr Ala Val Glu Lys Lys Glu Lys	
30 35 40	
cct ctt cca aga ctt aat atc cat tct gga ttc tgg att ttg gca tcc	196

Pro	Leu	Pro	Arg	Leu	Asn	Ile	His	Ser	Gly	Phe	Trp	Ile	Leu	Ala	Ser	
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att	gtt	gtg	acc	tat	tat	gtt	gac	ttc	ttt	aaa	acc	ctt	aaa	gaa	aac	244
Ile	Val	Val	Thr	Tyr	Tyr	Val	Asp	Phe	Phe	Lys	Thr	Leu	Lys	Glu	Asn	
60					65					70					75	
ttc	cac	act	agc	agc	tgg	ttt	ctc	tgt	ggc	agt	gcc	ttg	ttg	ctt	gtc	292
Phe	His	Thr	Ser	Ser	Trp	Phe	Leu	Cys	Gly	Ser	Ala	Leu	Leu	Leu	Val	
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agt	tta	tca	att	gca	ttt	tac	tgc	ata	gtc	tac	ctg	gaa	tgg	tat	tgt	340
Ser	Leu	Ser	Ile	Ala	Phe	Tyr	Cys	Ile	Val	Tyr	Leu	Glu	Trp	Tyr	Cys	
			95					100					105			
gga	att	gga	gaa	tat	gat	gtc	aag	tat	сса	gcc	ttg	ata	ccc	att	acc	388
Gly	Ile	Gly	Glu	Tyr	Asp	Val	Lys	Tyr	Pro	Ala	Leu	Ile	Pro	Ile	Thr	
		110					115					120				
act	gcc	tcc	ttt	att	gca	gca	gga	att	tgc	ttc	aac	att	gct	tta	tgg	436
Thr	Ala	Ser	Phe	Ile	Ala	Ala	Gly	Ile	Cys	Phe	Asn	Ile	Ala	Leu	Trp	
	125					130					135					
cat	gtg	tgg	tcg	ttt	ttc	act	cca	ttg	ttg	ttg	ttt	acc	cag	ttt	atg	484
His	Val	Trp	Ser	Phe	Phe	Thr	Pro	Leu	Leu	Leu	Phe	Thr	Gln	Phe	Met	
140					145					150					155	
ggg	gtt	gtc	atg	ttt	atc	aca	ctc	ctt	gga	tga	ttt	ccga	agag	ac		530
Gly	Val	Val	Met	Phe	Ile	Thr	Leu	Leu	Gly	į.						
				160)				165	,						
agg	gtct	tct	atgt	tgcc	ca g	gcte	tctt	t ga	acto	ctgg	gat	caag	gtga	tcct	cctgcc	590
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303/307

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aattttcaga	agtcagtgat	acagaagtac	tattttgcaa	tgttaatctg	tttgagtctt	890
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cactcttcca	tttaagggat	agcagttcct	tgtataaaat	gactggatgt	gtataaagga	1010
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agttttgtta	ggtgacagga	ccaaatgaaa	atattttatg	ttttctcatc	actttagatt	1130
ttatcattat	gtacattact	gggtttttag	catttcctaa	tgtgaagttt	taatcacttt	1190
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⟨220⟩

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cagccaggag cggtttctg ggaactgtgg gatgtgccct tgggggcccg agaaaacaga 180
aggaag atg ctc cag acc agt aac tac agc ctg gtg ctc tct ctg cag 228

Met Leu Gln Thr Ser Asn Tyr Ser Leu Val Leu Ser Leu Gln

5 10

ttc	ctg	ctg	ctg	tcc	tat	gac	ctc	ttt	gtc	aat	tcc	ttc	tca	gaa	ctg	276
Phe	Leu	Leu	Leų	Ser	Tyr	Asp	Leu	Phe	Val	Asn	Ser	Phe	Ser	Glu	Leu	
15			•		20				•	25					30	
ctc	caa	aag	act	cct	gtc	atc	cag	ctt	gtg	ctc	ttc	atc	atc	cag	gat	324
Leu	Gln	Lys	Thr	Pro	Val	Ile	Gln	Leu	Val	Leu	Phe	Ile	Ile	Gln	Asp	
				35					40					45		•
att	gca	gtc	ctc	ttc	aac	atc	atc	atc	att	ttc	ctc	atg	ttc	ttc	aac	372
Ile	Ala	Val	Leu	Phe	Asn	Ile	Ile	Ile	Ile	Phe	Leu	Met	Phe	Phe	Asn	
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acc	ttc	gtc	ttc	cag	gct	ggc	ctg	gtc	aac	ctc	cta	ttc	cat	aag	ttc	420
Thr	Phe	Val	Phe	G1n	Ala	Gly	Leu	Val	Asn	Leu	Leu	Phe	His	Lys	Phe	
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aaa	ggg	acc	atc	atc	ctg	aca	gct	gtg	tac	ttt	gcc	ctc	agc	atc	tcc	468
Lys	Gly	Thr	Ile	Ile	Leu	Thr	Ala	Val	Tyr	Phe	Ala	Leu	Ser	Ile	Ser	
	80					85					90					
ctt	cat	gtc	tgg	gtc	atg	aac	tta	cgc	tgg	aaa	aac	tcc	aac	agc	ttc	516
Leu	His	Val	Trp	Val	Met	Asn	Leu	Arg	Trp	Lys	Asn	Ser	Asn	Ser	Phe	
95					100					105					110	
ata	tgg	aca	gat	gga	ctt	caa	atg	ctg	ttt	gta	ttc	cag	aga	cta	gca	564
Ile	Trp	Thr	Asp	Gly	Leu	Gln	Met	Leu	Phe	Val	Phe	Gln	Arg	Leu	Ala	
				115					120					125		
gca	gtg	ttg	tac	tgc	tac	ttc	tat	aaa	cgg	aca	gcc	gta	aga	cta	ggc	612
Ala	Val	Leu	Tyr	Cys	Tyr	Phe	Tyr	Lys	Arg	Thr	Ala	Val	Arg	Leu	Gly	
			130					135					140)		
gat	cct	cac	ttc	tac	cag	gac	tct	ttg	tgg	ctg	cgc	aag	gag	tto	atg	660

305/307

Asp Pro His Phe Tyr Gln Asp Ser Leu Trp Leu Arg Lys Glu Phe Met

145 150 155

caa gtt cga agg tgacctct tgtcacactg atggatactt ttccttcctg 710

Gln Val Arg Arg

160

770 atagaagcca catttgctgc tttgcaggga gagttggccc tatgcatggg caaacagctg gactttccaa ggaaggttca gactagctgt gttcagcatt caagaaggaa gatcctccct 830 890 cttgcacaat tagagtgtcc ccatcggtct ccagtgcggc atcccttcct tgccttctac 950 ctctgttcca cccctttcc ttcctttcct ctctgtacca ttcattctcc ctgaccggcc 1010 tttcttgccg agggttctgt ggctcttacc cttgtgaagc ttttccttta gcctgggaca 1070 gaaggacctc ccagccccca aaggatctcc cagtgaccaa aggatgcgaa gagtgatagt 1130 tacgtgctcc tgactgatca caccgcagac atttagattt ttatacccaa ggcactttaa 1174 aaaaatgttt tataaataga gaataaattg aattcttgtt ccat 🕟

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<220>

<221> CDS

⟨222⟩ (208)... (873)

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cggcctccca gcgctcccaa gccgcagcgg ccgcgcccct tcagctagct cgctcgctcg 180

ctct	gctt	tcc (ctgct	tgcc	gg ci	tgcg	c a	tg g	cg ti	tg go	cg ti	tg g	cg g	cg ci	tg	231
							Mo	et A	la Le	eu Al	la Le	eu A	la A	la Ļ	eu .	
								1				5				
gcg	gcg	gtc	gag	ccg	gcc	tgc	ggc	agc	cgg	tac	cag	cag	ttg	cag	aat	279
Ala	Ala	Val	Glu	Pro	Ala	Cys	Gly	Ser	Arg	Tyr	Gln	Gln	Leu	Gln	Asn	
	10					15					20					
gaa	gaa	gag	tct	gga	gaa	cct	gaa	cag	gct	gca	ggt	gat	gct	cct	cca	327
Glu	Glu	G1u	Ser	Gly	Glu	Pro	Glu	Gln	Ala	Ala	Gly	Asp	Ala	Pro	Pro	
25					30					35					40	
cct	tac	agc	agc	att	tct	gca	gag	agc	gca	gca	tat	ttt	gac	tac	aag	375
Pro	Tyr	Ser	Ser	Ile	Ser	Ala	Glu	Ser	Ala	Ala	Tyr	Phe	Asp	Tyr	Lys	
			·	45					50	•				55		
gat	gag	tct	ggg	ttt	cca	aag	ccc	cca	tct	tac	aat	gta	gct	aca	aca	423
Asp	Glu	Ser	Gly	Phe	Pro	Lys	Pro	Pro	Ser	Tyr	Asn	Val	Ala	Thr	Thr	
			60					65					70			
ctg	ссс	agt	tat	gat	gaa	gcg	gag	agg	acc	aag	gct	gaa	gct	act	atc	471
Leu	Pro	Ser	Tyr	Asp	Glu	Ala	Glu	Arg	Thr	Lys	Ala	Glu	Ala	Thr	Ile	
		75					80					85				
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Pro	Leu	Val	Pro	Gly	Arg	Asp	Glu	Asp	Phe	Val	Gly	Arg	Asp	Asp	Phe	
	90					95					100					
gat	gat	gct	gac	cag	ctg	agg	ata	gga	aat	gat	ggg	att	ttc	atg	tta	567
Asp	Asp	Ala	Asp	Gln	Leu	Arg	Ile	Gly	Asn	Asp	Gly	Ile	Phe	Met	Leu	
105					110					115					120	
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Thr	Phe	Phe	Met	Ala	Phe	Leu	Phe	Asn	Trp	Ile	Gly	Phe	Phe	Leu	Ser	
				125					130				٠	135		
ttt	tgc	ctg	acc	act	tca	gct	gca	gga	agg	tat	ggg	gcc	att	tca	gga	663
Phe	Cys	Leu	Thr	Thr	Ser	Ala	Ala	Gly	Arg	Tyr	Gly	Ala	Ile	Ser	Gly	
			140					145					150			
ttt	ggt	ctc	tct	cta	att	aaa	tgg	atc	ctg	att	gtc	agg	ttt	tcc	acc	711
Phe	Gly	Leu	Ser	Leu	Ile	Lys	Trp	Ile	Leu	Ile	Val	Arg	Phe	Ser	Thr	
		155					160					165				
tat	ttc	cct	gga	tat	ttt	gat	ggt	cag	tac	tgg	ctc	tgg	tgg	gtg	ttc	759
Tyr	Phe	Pro	Gly	Tyr	Phe	Asp	Gly	Gln	Tyr	Trp	Leu	Trp	Trp	Val	Phe	
	170					175					180					
ctt	gtt	tta	ggc	ttt	ctc	ctg	ttt	ctc	aga	gga	ttt	atc	aat	tat	gca	807
Leu	Val	Leu	Gly	Phe	Leu	Leu	Phe	Leu	Arg	Gly	Phe	Ile	Asn	Tyr	Ala	
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Lys	Val	Arg	Lys	Met	Pro	Glu	Thr	Phe	Ser	Asn	Leu	Pro	Arg	Thr	Arg	
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gtt	ctc	ttt	att	tat	taaa	agat	gtt	ttctį	ggca	aa g	gccti	tcct	g ca	ttta	tgaa	910
Val	Leu	Phe	Île	Tyr			•									
			220													
ttc	tctc	tca a	agaa	gcaa	ga ga	aaca	cctg	cag	gaag	tgaa	tca	agat	gca	gaaca	acagag	970
gaa	taato	cac (ctgc	ttta	aa aa	aaata	aaag	t ac	tgtt	gaaa	ag					1012

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